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(54) Title: RG NUCLEIC ACIDS FOR CONFERRING DISEASE RESISTANCE TO PLANTS

(57) Abstract

The present invention provides RG nucleic acids and proteins which confer disease resistance to plants. The nucleic acids can be used to produce transgenic plants resistant to pests. Antibodies to proteins of the invention are also provided.

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RG NUCLEIC ACIDS FOR CONFERRING DISEASE RESISTANCE TO PLANTS

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The present application is a continuation-in-part application ("CIP") of U.S. Patent Application Serial No. ("USSN") 08/781,734, filed January 10, 1997. The 10 aforementioned application is explicitly incorporated herein by reference in its entirety and for all purposes.

This invention was made with Government support under Grant Nos. 92-37300-7547 and 95-37300-1571, awarded by the United States Department of Agriculture. 15 The Government has certain rights in this invention.

FIELD OF THE INVENTION

The present invention relates generally to plant molecular biology. In particular, it relates to nucleic acids and methods for conferring pest resistance in plants. 20 particularly lettuce.

BACKGROUND OF THE INVENTION

Recently, several resistance genes have been cloned by several groups from several plants. Many of these genes are sequence related. The derived amino acid 25 sequences of the most common class, *RPS2*, *RPM1* (bacterial resistances in *Arabidopsis* (Mindrinos *et al.* *Cell* 78:1089-1099 (1994)); Bent *et al.* *Science* 265:1856-1860 (1994); Grant *et al.*, *Science* 269:843-846 (1995)), *L6* (fungal resistance in flax; Lawrence, *et al.*, *The Plant Cell* 7:1195-1206 (1995)), and *N*, (virus resistance in tobacco; Whitham, *et al.*, *Cell* 78:1101-1115 (1994); and U.S. Patent No. 5,571,706), all contain leucine-rich 30 repeats (LRR) and nucleotide binding sites (NBS).

The NBS is a common motif in several mammalian gene families encoding signal transduction components (e.g., *Ras*) and is associated with ATP/GTP-binding sites.

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LRR domains can mediate protein-protein interactions and are found in a variety of proteins involved in signal transduction, cell adhesion and various other 5 functions. LRRs are leucine rich regions often comprising 20-30 amino acid repeats where leucine and other aliphatic residues occur periodically. LRRs can function extracellularly or intracellularly.

Since the onset of civilization, plant diseases have had catastrophic effects 10 on crops and the well-being of the human population. Plant diseases continue to effect enormous human and economic costs. An increasing human population and decreasing amounts of arable land make all approaches to preventing and treating plant pathogen destruction critical. The ability to control and enhance a plant's protective responses against pathogens would be of enormous benefit. Tissue-specific and temporal control of mechanisms responsible for plant cell death would also be of great practical and economic 15 value. The present invention fulfills these and other needs.

What is needed in the art are plant disease resistance genes and means to 20 create transgenic disease resistance plants, particularly in lettuce. Further, what is needed in the art is a means to DNA fingerprint cultivars and germplasm with respect to their disease resistance haplotypes for use in plant breeding programs. The present invention provides these and other advantages.

SUMMARY OF THE INVENTION

The present invention provides isolated nucleic acid constructs. These constructs comprise an RG (resistance gene) polynucleotide which encodes an RG 25 polypeptide having at least 60% sequence identity to an RG polypeptide selected from the group consisting of: an RG1 polypeptide, an RG2 polypeptide, an RG3 polypeptide, and an RG4 polypeptide. RG1, RG2, RG3, RG4, and the like, represent individual "RG families." Each "RG family," as defined herein, is a group of polypeptide sequences that have at least 60% amino acid sequence identity. Individual members of an RG family, 30 *i.e.*, individual species of the genus, typically map to the same genomic locus. The invention provides for constructs comprising nucleotides encoding the RG families of the

invention, which can include sequences encoding a leucine rich region (LRR), and/or a nucleotide binding site (NBS), or both.

The invention provides for an isolated nucleic acid construct comprising an RG polynucleotide which encodes an RG polypeptide having at least 60% sequence identity to an RG polypeptide from an RG family selected from the group consisting of: an RG1 polypeptide, an RG2 polypeptide, an RG3 polypeptide, an RG4 polypeptide, an RG5 polypeptide, and an RG7 polypeptide. In alternative embodiments, the nucleic acid construct comprises an RG polynucleotide which encodes an RG polypeptide comprising an leucine rich region (LRR), or, an RG polypeptide comprising a nucleotide binding site (NBS). The nucleic acid construct can comprise a polynucleotide which is a full length gene. In another embodiment, the nucleic acid construct encodes a fusion protein.

In one embodiment, the nucleic acid construct comprises a sequence encoding an RG1 polypeptide. The RG1 polypeptide can be encoded by a polynucleotide sequence selected from the group consisting of SEQ ID NO:1 (RG1A), SEQ ID NO:2 and SEQ ID NO:137 (RG1B), SEQ ID NO: 3 (RG1C), SEQ ID NO:4 (RG1D), SEQ ID NO:5 (RG1E), SEQ ID NO:6 (RG1F), SEQ ID NO:7 (RG1G), SEQ ID NO:8 (RG1H), SEQ ID NO:9 (RG1I), and SEQ ID NO:10 (RG1J).

In another embodiment, the nucleic acid construct comprises a sequence encoding an RG2 polypeptide. The RG2 polypeptide can be encoded by a polynucleotide sequence selected from the group consisting of: SEQ ID NO:21 and SEQ ID NO:27 (RG2A); SEQ ID NO:23 and SEQ ID NO:28 (RG2B); SEQ ID NO:29 (RG2C); SEQ ID NO:30 (RG2D); SEQ ID NO:31 (RG2E); SEQ ID NO:32 (RG2F); SEQ ID NO:33 (RG2G); SEQ ID NO:34 (RG2H); SEQ ID NO:35 (RG2I); SEQ ID NO:36 (RG2J); SEQ ID NO:37 (RG2K); SEQ ID NO:38 (RG2L); SEQ ID NO:39 (RG2M); SEQ ID NO:87 (RG2A); SEQ ID NO:89 (RG2B); SEQ ID NO:91 (RG2C); SEQ ID NO:93 (RG2D) and SEQ ID NO:94 (RG2D); SEQ ID NO:96 (RG2E); SEQ ID NO:98 (RG2F); SEQ ID NO:100 (RG2G); SEQ ID NO:102 (RG2H); SEQ ID NO:104 (RG2I); SEQ ID NO:106 (RG2J) and SEQ ID NO:107 (RG2J); SEQ ID NO:109 (RG2K) and (SEQ ID NO:110 (RG2K); SEQ ID NO:112 (RG2L); SEQ ID NO:114 (RG2M); SEQ ID NO:116 (RG2N); SEQ ID NO:118 (RG2O); SEQ ID NO:120 (RG2P); SEQ ID NO:122 (RG2Q); SEQ ID NO:124 (RG2S); SEQ ID NO:126 (RG2T); SEQ ID NO:128 (RG2U); SEQ ID NO:130 (RG2V); and, SEQ ID NO:132 (RG2W).

In other embodiments, the nucleic acid construct comprises a RG3 sequence (SEQ ID NO:68) encoding an RG3 polypeptide (SEQ ID NO:138) (RG3). In other embodiments, the nucleic acid construct comprises an RG4 sequence (SEQ ID NO:69) encoding an RG4 polypeptide (SEQ ID NO:139) (RG4).

5 In other embodiments, the nucleic acid construct comprises a RG5 sequence (SEQ ID NO:134) encoding an RG5 polypeptide (SEQ ID NO:135). The RG5 polypeptide can be encoded by a polynucleotide sequence as set forth in SEQ ID NO:134.

10 The invention also provides for a nucleic acid construct which comprises an RG7 sequence encoding an RG7 polypeptide. The RG7 polypeptide can be encoded by a polynucleotide sequence as set forth in SEQ ID NO:136.

15 In further embodiments, the nucleic acid construct can further comprise a promoter operably linked to the RG polynucleotide. In alternative embodiments, the promoter can be a plant promoter; a disease resistance promoter; a lettuce promoter; a constitutive promoter; an inducible promoter; or, a tissue-specific promoter. The nucleic acid construct can comprise a promoter sequence from an RG gene linked to a heterologous polynucleotide.

20 The invention also provides for a transgenic plant comprising a recombinant expression cassette comprising a promoter operably linked to an RG polynucleotide. The expression cassette can comprise a plant promoter or a viral promoter; the plant promoter can be a heterologous promoter. In one embodiment, the transgenic plant is lettuce. In alternative embodiments, the transgenic plant comprises an expression cassette which includes an RG polynucleotide selected from the group consisting of SEQ ID NO:1 (RG1A); SEQ ID NO:2 and SEQ ID NO:137 (RG1B); SEQ ID NO:3 (RG1C); SEQ ID NO:4 (RG1D); SEQ ID NO:5 (RG1E); SEQ ID NO:6 (RG1F); SEQ ID NO:7 (RG1G);
25 SEQ ID NO:8 (RG1H); SEQ ID NO:9 (RG1I) and SEQ ID NO:10 (RG1J); SEQ ID NO:21 and SEQ ID NO:27 (RG2A); SEQ ID NO:23 and SEQ ID NO:28 (RG2B); SEQ ID NO:29 (RG2C); SEQ ID NO:30 (RG2D); SEQ ID NO:31 (RG2E); SEQ ID NO:32 (RG2F); SEQ ID NO:33 (RG2G); SEQ ID NO:34 (RG2H); SEQ ID NO:35 (RG2I); SEQ ID NO:36 (RG2J); SEQ ID NO:37 (RG2K); SEQ ID NO:38 (RG2L); SEQ ID NO:39 (RG2M); SEQ ID NO:87 (RG2A); SEQ ID NO:89 (RG2B); SEQ ID NO:91 (RG2C); SEQ ID NO:93 (RG2D) and SEQ ID NO:94 (RG2D); SEQ ID NO:96 (RG2E); SEQ ID NO:98 (RG2F); SEQ ID NO:100 (RG2G); SEQ ID NO:102 (RG2H); SEQ ID NO:104

(RG2I); SEQ ID NO:106 (RG2J) and SEQ ID NO:107 (RG2J); SEQ ID NO:109 (RG2K) and (SEQ ID NO:110 (RG2K); SEQ ID NO:112 (RG2L); SEQ ID NO:114 (RG2M); SEQ ID NO:116 (RG2N); SEQ ID NO:118 (RG2O); SEQ ID NO:120 (RG2P); SEQ ID NO:122 (RG2Q); SEQ ID NO:124 (RG2S); SEQ ID NO:126 (RG2T); SEQ ID NO:128 (RG2U); SEQ ID NO:130 (RG2V); and, SEQ ID NO:132 (RG2W); SEQ ID NO:68 (RG3); SEQ ID NO:69 (RG4); SEQ ID NO:134 (RG5); or SEQ ID NO:136 (RG7).

The invention provide for a transgenic plant comprising an expression cassette comprising an RG polynucleotide which can encode an RG1 polypeptide selected from the group consisting of SEQ ID NO:11 (RG1A), SEQ ID NO:12 (RG1B), SEQ ID NO:13 (RG1C), SEQ ID NO:14 (RG1D), SEQ ID NO:15 (RG1E), SEQ ID NO:16 (RG1F), SEQ ID NO:17 (RG1G), SEQ ID NO:18 (RG1H), SEQ ID NO:19 (RG1I), or SEQ ID NO:20 (RG1J); or, an RG2 polypeptide selected from the group consisting of SEQ ID NO:22 and SEQ ID NO:41 (RG2A); SEQ ID NO:24 and SEQ ID NO:42 (RG2B); SEQ ID NO:43 (RG2C); SEQ ID NO:44 (RG2D); SEQ ID NO:45 (RG2E); SEQ ID NO:46 (RG2F); SEQ ID NO:47 (RG2G); SEQ ID NO:48 (RG2H); SEQ ID NO:49 (RG2I); SEQ ID NO:50 (RG2J); SEQ ID NO:51 (RG2K); SEQ ID NO:52 (RG2L); SEQ ID NO:53 (RG2M); SEQ ID NO:88 (RG2A); SEQ ID NO:90 (RG2B); SEQ ID NO:92 (RG2C); SEQ ID NO:95 (RG2D); SEQ ID NO:97 (RG2E); SEQ ID NO:99 (RG2F); SEQ ID NO:101 (RG2G); SEQ ID NO:103 (RG2H); SEQ ID NO:105 (RG2I); SEQ ID NO:108 (RG2J); SEQ ID NO:111 (RG2K); SEQ ID NO:113 (RG2L); SEQ ID NO:115 (RG2M); SEQ ID NO:117 (RG2N); SEQ ID NO:119 (RG2O); SEQ ID NO:121 (RG2P); SEQ ID NO:123 (RG2Q); SEQ ID NO:125 (RG2S); SEQ ID NO:127 (RG2T); SEQ ID NO:129 (RG2U); SEQ ID NO:131 (RG2V); and, SEQ ID NO:133 (RG2W); an RG4 polypeptide as set forth by SEQ ID NO:72; an RG5 polypeptide with a sequence as set forth by SEQ ID NO:135; or, an RG7 polypeptide.

The invention also provides for a method of enhancing disease resistance in a plant, the method comprising introducing into the plant a recombinant expression cassette comprising a promoter functional in the plant and operably linked to an RG polynucleotide sequence. In this method, the plant can be a lettuce plant; and, the RG polynucleotide can encode an RG polypeptide selected from the group consisting of an RG1 polypeptide selected from the group consisting of SEQ ID NO:11 (RG1A), SEQ ID NO:12 (RG1B), SEQ ID NO:13 (RG1C), SEQ ID NO:14 (RG1D), SEQ ID NO:15 (RG1E), SEQ ID

NO:16 (RG1F), SEQ ID NO:17 (RG1G), SEQ ID NO:18 (RG1H), SEQ ID NO:19 (RG1I), or SEQ ID NO:20 (RG1J); or, an RG2 polypeptide selected from the group consisting of SEQ ID NO:22 and SEQ ID NO:41 (RG2A); SEQ ID NO:24 and SEQ ID NO:42 (RG2B); SEQ ID NO:43 (RG2C); SEQ ID NO:44 (RG2D); SEQ ID NO:45 (RG2E); SEQ ID NO:46 (RG2F); SEQ ID NO:47 (RG2G); SEQ ID NO:48 (RG2H); SEQ ID NO:49 (RG2I); SEQ ID NO:50 (RG2J); SEQ ID NO:51 (RG2K); SEQ ID NO:52 (RG2L); SEQ ID NO:53 (RG2M); SEQ ID NO:72; SEQ ID NO:74; SEQ ID NO:88 (RG2A); SEQ ID NO:90 (RG2B); SEQ ID NO:92 (RG2C); SEQ ID NO:95 (RG2D); SEQ ID NO:97 (RG2E); SEQ ID NO:99 (RG2F); SEQ ID NO:101 (RG2G); SEQ ID NO:103 (RG2H); SEQ ID NO:105 (RG2I); SEQ ID NO:108 (RG2J); SEQ ID NO:111 (RG2K); SEQ ID NO:113 (RG2L); SEQ ID NO:115 (RG2M); SEQ ID NO:117 (RG2N); SEQ ID NO:119 (RG2O); SEQ ID NO:121 (RG2P); SEQ ID NO:123 (RG2Q); SEQ ID NO:125 (RG2S); SEQ ID NO:127 (RG2T); SEQ ID NO:129 (RG2U); SEQ ID NO:131 (RG2V); and, SEQ ID NO:133 (RG2W). In this method, the promoter can be a plant disease resistance promoter, a tissue-specific promoter, a constitutive promoter, or an inducible promoter.

The invention also provides for a method of detecting RG resistance genes in a nucleic acid sample, the method comprising: contacting the nucleic acid sample with an RG polynucleotide to form a hybridization complex; and, wherein the formation of the hybridization complex is used to detect the RG resistance gene in the nucleic acid sample. In this method, the RG polynucleotide can be an RG1 polynucleotide, an RG2 polynucleotide, an RG3 polynucleotide, an RG4 polynucleotide, an RG5 polynucleotide or an RG7 polynucleotide. In this method, the RG resistance gene can be amplified prior to the step of contacting the nucleic acid sample with the RG polynucleotide, and, the RG resistance gene can be amplified by the polymerase chain reaction. In one embodiment, the RG polynucleotide is labeled.

The invention further provides for an RG polypeptide having at least 60% sequence identity to a polypeptide selected from the group consisting of: an RG1 polypeptide, an RG2 polypeptide, an RG3 polypeptide, an RG4 polypeptide, an RG5 polypeptide, and an RG7 polypeptide.

A further understanding of the nature and advantages of the present invention may be realized by reference to the remaining portions of the specification, the figures and claims.

All publications, patents and patent applications cited herein are hereby
5 expressly incorporated by reference for all purposes.

DETAILED DESCRIPTION OF THE INVENTION

This invention relates to families of RG genes, particularly from *Lactuca sativa*. Nucleic acid sequences of the present invention can be used to confer resistance in
10 plants to a variety of pests including viruses, fungi, nematodes, insects, and bacteria. Sequences from within the RG genes can be used to fingerprint cultivars or germplasm for the presence of desired resistance genes. Promoters of RG genes can be used to drive heterologous gene expression under conditions in which RG genes are expressed. Further,
15 the present invention provides RG proteins and antibodies specifically reactive to RG proteins. Antibodies to RG proteins can be used to detect the type and amount of RG protein expressed in a plant sample.

The present invention has use over a broad range of types of plants, including species from the genera *Cucurbita*, *Rosa*, *Vitis*, *Juglans*, *Fragaria*, *Lotus*,
20 *Medicago*, *Onobrychis*, *Trifolium*, *Trigonella*, *Vigna*, *Citrus*, *Linum*, *Geranium*, *Manihot*,
Daucus, *Arabidopsis*, *Brassica*, *Raphanus*, *Sinapis*, *Atropa*, *Capsicum*, *Datura*,
Hyoscyamus, *Lycopersicon*, *Nicotiana*, *Solanum*, *Petunia*, *Digitalis*, *Majorana*,
25 *Ciahorium*, *Helianthus*, *Lactuca*, *Bromus*, *Asparagus*, *Antirrhinum*, *Heterocallis*, *Nemesis*,
Pelargonium, *Panieum*, *Pennisetum*, *Ranunculus*, *Senecio*, *Salpiglossis*, *Cucumis*,
Browalia, *Glycine*, *Pisum*, *Phaseolus*, *Lolium*, *Oryza*, *Zea*, *Avena*, *Hordeum*, *Secale*,
30 *Triticum*, and, *Sorghum*. In particularly preferred embodiments, species from the family
Compositae and in particular the genus *Lactuca* are employed such as *L. sativa* and such
subspecies as *crispula*, *longifolia*, and *asparagina*.

The nucleic acids of the present invention can be used in marker-aided selection. Marker-aided selection does not require the complete sequence of the gene or
30 precise knowledge of which sequence confers which specificity. Instead, partial sequences can be used as hybridization probes or as the basis for oligonucleotide primers to amplify nucleic acid, e.g., by PCR. Partial sequences can be used in other methods, such as to

follow the segregation of chromosome segments containing resistance genes in plants. Because the RG marker is the gene itself, there can be negligible recombination between the marker and the resistance phenotype. Thus, RG polynucleotides of the present invention provide an optimal means to DNA fingerprint cultivars and wild germplasm with respect to their disease resistance haplotypes. This can be used to indicate which 5 germplasm accessions and cultivars carry the same resistance genes. At present, selection of plants (e.g., lettuce) for resistance to some diseases is slow and difficult. But linked markers allow indirect selection for such resistance genes. Moreover, RG markers also allow resistance genes to be identified and combined in a manner that would not otherwise 10 be possible. Numerous accessions have been identified that provide resistance to all isolates of downy mildew (*Bremia lactucae*). However, without molecular markers it is impossible to combine such resistances from different sources. The nucleic acid sequences of the invention provide for a fast and convenient means to identify and combine 15 resistances from different sources. The RG markers of the invention can also be used to identify recombinants that have new combinations of resistance genes in *cis* on the same chromosome.

In addition, RG markers may allow the identification of the Mendelian factors determining traits, such as field resistance to downy mildew. Once such markers have been identified, they will greatly increase the ease with which field resistance can be 20 transferred between lines and combined with other resistances.

In another application, primers to RG sequences can be also designed to amplify sequences that are conserved in multiple RG family members. This gives genetic information on multiple RG family members. Alternatively, one or more primers can be made to sequences unique to a single resistance gene genus or a single RG specie. This 25 allows an analysis of individual family groups (an RG genus) or an individual family member (a specie). Primers made to individual RGs at the edge of each cluster can be used to select for recombinants within the cluster. This minimizes the amount of linkage drag during introgression. Classical and molecular genetics has shown that pest resistance genes tend to be clustered in the genome. Pest resistance loci comprise arrays of genes 30 and exhibit a variety of complex haplotypes rather than being simple alternate allelic forms. Pest resistance is conferred by families, or genera, of related RG sequences, individual members, or species, of which have evolved to have a different specificity.

Oligonucleotide primers can be designed that amplify members from multiple haplotypes, or genuses, or amplify only members of one genus, or only amplify an individual specie. This will provide codominant information and allow heterozygotes to be distinguished from homozygotes.

5 Further, comparison of RG sequences will allow a determination of which sequences are critical for resistance and will ultimately lead to engineering resistance genes with new specificities. Resistance gene sequences were not previously available for lettuce. Marker-aided selection will greatly increase the precision and speed of breeding for disease resistance. Transgenic approaches will allow pyramiding of resistance genes
10 into a single Mendelian unit, transfer between sexually-incompatible species, substitute for conventional backcrossing procedures, and allow expression of other genes in parallel with resistance genes.

The RG polynucleotides also have utility in the construction of disease resistant transgenic plants. This avoids lengthy and sometimes difficult backcrossing
15 programs currently necessary for introgression of resistance. It is also possible to transfer resistance polynucleotides between sexually-incompatible species, thereby greatly increasing the germplasm pool that can be used as a source of resistance genes. Cloning of multiple RG sequences in a single cassette will allow pyramiding of genes for resistance against multiple isolates of a single pathogen such as downy mildew or against multiple
20 pathogens. Once introduced, such a cassette can be manipulated by classical breeding methods as a single Mendelian unit.

Transgenic plants of the present invention can also be constructed using an RG promoter. The promoter sequences from RG sequences of the invention can be used with RG genes or heterologous genes. Thus, RG promoters can be used to express a
25 variety of genes in the same temporal and spatial patterns and at similar levels to resistance genes.

Nucleic acids of the Invention and Their Preparation

RG Polynucleotide Families

30 The present invention provides isolated nucleic acid constructs which comprise an RG polynucleotide. In alternative embodiments, the RG polynucleotide is at least 18 nucleotides in length, typically at least 20, 25, or 30 nucleotides in length, more

typically at least 100 nucleotides in length, generally at least 200 nucleotides in length, preferably at least 300 nucleotides in length, more preferably at least 400 nucleotides in length, and most preferably at least 500 nucleotides in length.

In particularly preferred embodiments, the RG polynucleotide encodes a RG protein which confers resistance to plant pests. This RG protein can be longer, equivalent, or shorter than the RG protein encoded by an RG gene. In various embodiments, an RG polynucleotide can hybridize under stringent conditions to members of an RG family (an RG genus); *e.g.*, it can hybridize to a member of the RG1 RG family, such as an RG1 polynucleotide selected from the group consisting of: SEQ ID NO:1 (RG1A); SEQ ID NO:2 and SEQ ID NO:137 (RG1B); SEQ ID NO: 3 (RG1C); SEQ ID NO:4 (RG1D); SEQ ID NO:5 (RG1E); SEQ ID NO:6 (RG1F); SEQ ID NO:7 (RG1G); SEQ ID NO:8 (RG1H); SEQ ID NO:9 (RG1I) and SEQ ID NO:10 (RG1J).

In other embodiments, the polynucleotide can also hybridize under stringent conditions to a member of the RG2 family; such as an RG2 polynucleotide selected from the group consisting of: SEQ ID NO:21 and SEQ ID NO:27 (RG2A); SEQ ID NO:23 and SEQ ID NO:28 (RG2B); SEQ ID NO:29 (RG2C); SEQ ID NO:30 (RG2D); SEQ ID NO:31 (RG2E); SEQ ID NO:32 (RG2F); SEQ ID NO:33 (RG2G); SEQ ID NO:34 (RG2H); SEQ ID NO:35 (RG2I); SEQ ID NO:36 (RG2J); SEQ ID NO:37 (RG2K); SEQ ID NO:38 (RG2L); SEQ ID NO:39 (RG2M); SEQ ID NO:87 (RG2A); SEQ ID NO:89 (RG2B); SEQ ID NO:91 (RG2C); SEQ ID NO:93 (RG2D) and SEQ ID NO:94 (RG2D); SEQ ID NO:96 (RG2E); SEQ ID NO:98 (RG2F); SEQ ID NO:100 (RG2G); SEQ ID NO:102 (RG2H); SEQ ID NO:104 (RG2I); SEQ ID NO:106 (RG2J) and SEQ ID NO:107 (RG2J); SEQ ID NO:109 (RG2K) and (SEQ ID NO:110 (RG2K); SEQ ID NO:112 (RG2L); SEQ ID NO:114 (RG2M); SEQ ID NO:116 (RG2N); SEQ ID NO:118 (RG2O); SEQ ID NO:120 (RG2P); SEQ ID NO:122 (RG2Q); SEQ ID NO:124 (RG2S); SEQ ID NO:126 (RG2T); SEQ ID NO:128 (RG2U); SEQ ID NO:130 (RG2V); and, SEQ ID NO:132 (RG2W).

In alternative embodiments, each RG2 gene can also include an AC15 sequence which hybridizes under stringent conditions to a polynucleotide selected from the group consisting of: SEQ ID NO:56 (AC15-2A); SEQ ID NO:57 (AC15-2B); SEQ ID NO:58 (AC15-2C); SEQ ID NO:59 (AC15-2D); SEQ ID NO:60 (AC15-2E); SEQ ID NO:61 (AC15-2G); SEQ ID NO:62 (AC15-2H); SEQ ID NO:63 (AC15-2I); SEQ ID

NO:64 (AC15-2J); SEQ ID NO:65 (AC15-2L); SEQ ID NO:66 (AC15-2N); SEQ ID NO:67 (AC15-2O).

In other embodiments, an RG polynucleotide can hybridize under stringent conditions to an RG3 (SEQ ID NO:68), an RG4 (SEQ ID NO:69), and RG5 (SEQ ID NO:135), and an RG7 (SEQ ID NO:137), RG family member.

The present invention further provides nucleic acid constructs which comprise an RG polynucleotide which encodes RG polypeptides from various RG families; such as an RG polypeptide having at least 60% sequence identity to an RG polypeptide selected from the group consisting of: an RG1 polypeptide, an RG2 polypeptide, an RG3 polypeptide, and RG4 polypeptide, and RG5 polypeptide, and an RG7 polypeptide.

Exemplary RG1 polypeptides have the sequences shown in SEQ ID NO:2 (RG1A), SEQ ID NO:4 (RG1B), SEQ ID NO:6 (RG1C), SEQ ID NO:8 (RG1D), SEQ ID NO:10 (RG1E), SEQ ID NO:12 (RG1F), SEQ ID NO:14 (RG1G), SEQ ID NO:16 (RG1H), SEQ ID NO:20 (RG1J). Exemplary RG2 polypeptides have the sequences shown in SEQ ID NO:22 and SEQ ID NO:41 (RG2A); SEQ ID NO:24 and SEQ ID NO:42 (RG2B); SEQ ID NO:43 (RG2C); SEQ ID NO:44 (RG2D); SEQ ID NO:45 (RG2E); SEQ ID NO:46 (RG2F); SEQ ID NO:47 (RG2G); SEQ ID NO:48 (RG2H); SEQ ID NO:49 (RG2I); SEQ ID NO:50 (RG2J); SEQ ID NO:51 (RG2K); SEQ ID NO:52 (RG2L); SEQ ID NO:53 (RG2M); SEQ ID NO:88 (RG2A); SEQ ID NO:90 (RG2B); SEQ ID NO:92 (RG2C); SEQ ID NO:95 (RG2D); SEQ ID NO:97 (RG2E); SEQ ID NO:99 (RG2F); SEQ ID NO:101 (RG2G); SEQ ID NO:103 (RG2H); SEQ ID NO:105 (RG2I); SEQ ID NO:108 (RG2J); SEQ ID NO:111 (RG2K); SEQ ID NO:113 (RG2L); SEQ ID NO:115 (RG2M); SEQ ID NO:117 (RG2N); SEQ ID NO:119 (RG2O); SEQ ID NO:121 (RG2P); SEQ ID NO:123 (RG2Q); SEQ ID NO:125 (RG2S); SEQ ID NO:127 (RG2T); SEQ ID NO:129 (RG2U); SEQ ID NO:131 (RG2V); and, SEQ ID NO:133 (RG2W).

An exemplary RG3 polypeptide has the sequence shown in SEQ ID NO:138. An exemplary RG4 polypeptide has the sequence shown in SEQ ID NO:139. RG polynucleotides will have at least 60% identity, more typically at least 65% identity, generally at least 70% identity, and preferably at least 75% identity, more preferably at least 80% identity, and most preferably at least 85%, 90%, or 95% identity at the deduced amino acid level. The regions where substantial identity is assessed can be inclusive or exclusive of the nucleotide binding site or the leucine rich region.

Vectors and Transcriptional Control Elements

The invention, providing methods and reagents for making novel species and genuses of RG nucleic acids described herein, further provides methods and reagents for expressing these nucleic acids using novel expression cassettes, vectors, transgenic plants and animals, using constitutive and inducible transcriptional and translational *cis*- (e.g., promoters and enhancers) and *trans*-acting control elements.

The expression of natural, recombinant or synthetic plant disease resistance polypeptide-encoding or other (*i.e.*, antisense, ribozyme) nucleic acids can be achieved by operably linking the coding region a promoter (that can be plant-specific or not, 10 constitutive or inducible), incorporating the construct into an expression cassette (such as an expression vector), and introducing the resultant construct into an *in vitro* reaction system or a suitable host cell or organism. Synthetic procedures may also be used. Typical expression systems contain, in addition to coding or antisense sequence, 15 transcription and translation terminators, polyadenylation sequences, transcription and translation initiation sequences, and promoters useful for transcribing DNA into RNA. The expression systems optionally at least one independent terminator sequence, sequences permitting replication of the cassette *in vivo*, *e.g.*, plants, eukaryotes, or prokaryotes, or a combination thereof, (*e.g.*, shuttle vectors) and selection markers for the selected expression system, *e.g.*, plant, prokaryotic or eukaryotic systems. To ensure proper 20 polypeptide expression under varying conditions, a polyadenylation region at the 3'-end of the coding region can be included (see Li (1997) *Plant Physiol.* 115:321-325, for a review of the polyadenylation of RNA in plants). The polyadenylation region can be derived from the natural gene, from a variety of other plant genes, or from T-DNA (*e.g.*, using *Agrobacterium tumefaciens* T-DNA replacement vectors, see *e.g.*, Thykjaer (1997) *Plant Mol Biol.* 35:523-530; using a plasmid containing a gene of interest flanked by 25 *Agrobacterium* T-DNA border repeat sequences; Hansen (1997) "T-strand integration in maize protoplasts after codelivery of a T-DNA substrate and virulence genes," *Proc. Natl. Acad. Sci. USA* 94:11726-11730).

To identify the promoters, the 5' portions of the clones described here are 30 analyzed for sequences characteristic of promoter sequences. For instance, promoter sequence elements include the TATA box consensus sequence (TATAAT), which is usually 20 to 30 base pairs upstream of the transcription start site. In plants, further

upstream from the TATA box, at positions -80 to -100, there is typically a promoter element with a series of adenines surrounding the trinucleotide G (or T) N G (see, e.g., Messing, in *Genetic Engineering in Plants*, pp. 221-227, Kosage, Meredith and Hollaender, eds. 1983). If proper polypeptide expression is desired, a polyadenylation region at the 3'-end of the RG coding region should be included. The polyadenylation region can be derived from the natural gene, from a variety of other plant genes, or from viral genes, such as T-DNA.

The nucleic acids of the invention can be expressed in expression cassettes, vectors or viruses which are transiently expressed in cells using, for example, episomal expression systems (e.g., cauliflower mosaic virus (CaMV) viral RNA is generated in the nucleus by transcription of an episomal minichromosome containing supercoiled DNA, Covey (1990) *Proc. Natl. Acad. Sci. USA* 87:1633-1637). Alternatively, coding sequences can be inserted into the host cell genome becoming an integral part of the host chromosomal DNA.

Selection markers can be incorporated into expression cassettes and vectors to confer a selectable phenotype on transformed cells and sequences coding for episomal maintenance and replication such that integration into the host genome is not required. For example, the marker may encode biocide resistance, such as antibiotic resistance, particularly resistance to chloramphenicol, kanamycin, G418, bleomycin, hygromycin, or herbicide resistance, such as resistance to chlorosulfuron or Basta, to permit selection of those cells transformed with the desired DNA sequences, see for example, Blondelet-Rouault (1997) *Gene* 190:315-317; Aubrecht (1997) *J. Pharmacol. Exp. Ther.* 281:992-997. Because selectable marker genes conferring resistance to substrates like neomycin or hygromycin can only be utilized in tissue culture, chemoresistance genes are also used as selectable markers *in vitro* and *in vivo*. See also, Mengiste (1997) "High-efficiency transformation of *Arabidopsis thaliana* with a selectable marker gene regulated by the T-DNA 1' promoter," *Plant J.* 12:945-948, showing that the 1' promoter is an attractive alternative to the cauliflower mosaic virus (CaMV) 35S promoter for the generation of T-DNA insertion lines, the 1' promoter may be especially beneficial for the secondary transformation of transgenic strains containing the 35S promoter to exclude homology-mediated gene silencing.

The endogenous promoters from the RG genes of the present invention can be used to direct expression of the genes. These promoters can also be used to direct expression of heterologous structural genes. The promoters can be used, for example, in recombinant expression cassettes to drive expression of genes conferring resistance to any number of pathogens or pests, including fungi, bacteria, and the like.

Constitutive Promoters

In construction of recombinant expression cassettes, vectors, transgenics, of the invention, a promoter fragment can be employed to direct expression of the desired gene in all tissues of a plant or animal. Promoters that drive expression continuously under physiological conditions are referred to as "constitutive" promoters and are active under most environmental conditions and states of development or cell differentiation. Examples of constitutive promoters include those from viruses which infect plants, such as the cauliflower mosaic virus (CaMV) 35S transcription initiation region; the 1'- or 2'- promoter derived from T-DNA of *Agrobacterium tumefaciens*; the promoter of the tobacco mosaic virus; and, other transcription initiation regions from various plant genes known to those of skill. See also Holtorf (1995) "Comparison of different constitutive and inducible promoters for the overexpression of transgenes in *Arabidopsis thaliana*," *Plant Mol. Biol.* 29:637-646.

Inducible Promoters

Alternatively, a plant promoter may direct expression of the plant disease resistance nucleic acid of the invention under the influence of changing environmental conditions or developmental conditions. Examples of environmental conditions that may effect transcription by inducible promoters include pathogenic attack, anaerobic conditions, elevated temperature, drought, or the presence of light. Such promoters are referred to herein as "inducible" promoters. For example, the invention incorporates the drought-inducible promoter of maize (Busk (1997) *supra*); the cold, drought, and high salt inducible promoter from potato (Kirch (1997) *Plant Mol. Biol.* 33:897-909).

Embodiments of the invention also incorporate use of plant promoters which are inducible upon injury or infection to express the invention's plant disease resistance (RG) polypeptides. Various embodiments include use of, e.g., the promoter for a tobacco (*Nicotiana tabacum*) sesquiterpene cyclase gene (EAS4 promoter), which is expressed in wounded leaves, roots, and stem tissues, and upon infection with microbial pathogens (Yin

(1997) *Plant Physiol.* 115(2):437-451); the ORF13 promoter from *Agrobacterium rhizogenes* 8196, which is wound inducible in a limited area adjacent to the wound site (Hansen (1997) *Mol. Gen. Genet.* 254:337-343); the Shpx6b gene promoter, which is a plant peroxidase gene promoter induced by microbial pathogens (demonstrated using a 5 fungal pathogen, see Curtis (1997) *Mol. Plant Microbe Interact.* 10:326-338); the wound-inducible gene promoter wun1, derived from potato (Siebertz (1989) *Plant Cell* 1:961-968); the wound-inducible *Agrobacterium pmas* gene (mannopine synthesis gene) promoter (Guevara-Garcia (1993) *Plant J.* 4:495-505).

Alternatively, plant promoters which are inducible upon exposure to plant 10 hormones, such as auxins, are used to express the nucleic acids of the invention. For example, the invention can use the auxin-response elements E1 promoter fragment (AuxREs) in the soybean (*Glycine max L.*) (Liu (1997) *Plant Physiol.* 115:397-407); the auxin-responsive Arabidopsis GST6 promoter (also responsive to salicylic acid and hydrogen peroxide) (Chen (1996) *Plant J.* 10: 955-966); the auxin-inducible parC 15 promoter from tobacco (Sakai (1996) 37:906-913); a plant biotin response element (Streit (1997) *Mol. Plant Microbe Interact.* 10:933-937); and, the promoter responsive to the stress hormone abscisic acid (Sheen (1996) *Science* 274:1900-1902).

Plant promoters which are inducible upon exposure to chemicals reagents 20 which can be applied to the plant, such as herbicides or antibiotics, are also used to express the nucleic acids of the invention. For example, the maize In2-2 promoter, activated by benzenesulfonamide herbicide safeners, can be used (De Veylder (1997) *Plant Cell Physiol.* 38:568-577); application of different herbicide safeners induces distinct gene expression patterns, including expression in the root, hydathodes, and the shoot apical meristem. Coding sequence can be under the control of, e.g., a tetracycline-inducible 25 promoter, e.g., as described with transgenic tobacco plants containing the *Avena sativa L.* (oat) arginine decarboxylase gene (Masgrau (1997) *Plant J.* 11:465-473); or, a salicylic acid-responsive element (Stange (1997) *Plant J.* 11:1315-1324. Using chemically- (e.g., hormone- or pesticide-) induced promoters, harvesting of fruits and plant parts would be greatly facilitated. A chemical which can be applied to the transgenic plant in the field and 30 induce expression of a polypeptide of the invention throughout all or most of the plant would make a environmentally safe defoliant or herbicide. Thus, the invention also provides for transgenic plants containing an inducible gene encoding for the RG

polypeptides of the invention whose host range is limited to target plant species, such as weeds or crops before, during or after harvesting.

Abcission promoters are activated upon plant ripening, such as fruit ripening, and are especially useful incorporated in the expression systems (e.g., expression 5 cassettes, vectors) of the invention. In some embodiments, when a plant disease resistant polypeptide-encoding nucleic acid is under the control of such a promoter, rapid cell death, induced by expression of the invention's polypeptide, can accelerate and/or accentuate abcission, increasing the efficiency of the harvesting of fruits or other plant parts, such as cotton, and the like. Induction of rapid cell death at this time would accelerate separation 10 of the fruit from the plant, greatly augmenting harvesting procedures. See, e.g., Kalaitzis (1997) *Plant Physiol.* 113:1303-1308, discussing tomato leaf and flower abscission; Payton (1996) *Plant Mol. Biol.* 31:1227-1231, discussing ethylene receptor expression regulation during fruit ripening, flower senescence and abscission; Koehler (1996) *Plant Mol. Biol.* 31:595-606, discussing the gene promoter for a bean abscission cellulase; Kalaitzis (1995) 15 *Plant Mol. Biol.* 28: 647-656, discussing cloning of a tomato polygalacturonase expressed in abscission; del Campillo (1996) *Plant Physiol.* 111:813-820, discussing pedicel breakstrength and cellulase gene expression during tomato flower abscission.

Tissue-Specific Promoters

Tissue specific promoters are transcriptional control elements that are only 20 active in particular cells or tissues. Plant promoters which are active only in specific tissues or at specific times during plant development are used to express the nucleic acids of the invention. Examples of promoters under developmental control include promoters that initiate transcription only in certain tissues, such as leaves, roots, fruit, seeds, ovules, pollen, pistils, or flowers. Such promoters are referred to as "tissue specific". The 25 operation of a promoter may also vary depending on its location in the genome. Thus, an inducible promoter may become fully or partially constitutive in certain locations.

For example, a seed-specific promoter directs expression in seed tissues. Such promoters may be, for example, ovule-specific, embryo-specific, endosperm-specific, integument-specific, seed coat-specific, or some combination thereof. A leaf-specific 30 promoter has been identified in maize, Busk (1997) *Plant J.* 11:1285-1295. The ORF13 promoter from *Agrobacterium rhizogenes* exhibits high activity in roots (Hansen (1997) *supra*). A maize pollen-specific promoter has been identified in maize (Guerrero (1990)

Mol. Gen. Genet. 224:161-168). A tomato promoter active during fruit ripening, senescence and abscission of leaves and, to a lesser extent, of flowers can be used (Blume (1997) *Plant J.* 12:731-746). A pistol specific promoter has been identified in the potato (*Solanum tuberosum* L.) SK2 gene, encoding a pistil-specific basic endochitinase (Ficker (1997) *Plant Mol. Biol.* 35:425-431). The Blec4 gene from pea (*Pisum sativum* cv. 5 *Alaska*) is active in epidermal tissue of vegetative and floral shoot apices of transgenic alfalfa, making it a useful tool to target the expression of foreign genes to the epidermal layer of actively growing shoots. The activity of the Blec4 promoter in the epidermis of the shoot apex makes it particularly suitable for genetically engineering defense against insects 10 and diseases that attack the growing shoot apex (Mandaci (1997) *Plant Mol Biol.* 34:961-965).

The invention also provides for use of tissue-specific plant promoters include a promoter from the ovule-specific *BEL1* gene described in Reiser (1995) *Cell* 83:735-742, GenBank No. U39944. Suitable seed specific promoters are derived from the 15 following genes: *MAC1* from maize, Sheridan (1996) *Genetics* 142:1009-1020; *Cat3* from maize, GenBank No. L05934, Abler (1993) *Plant Mol. Biol.* 22:10131-1038; the gene encoding oleosin 18kD from maize, GenBank No. J05212, Lee (1994) *Plant Mol. Biol.* 26:1981-1987; viviparous-1 from *Arabidopsis*, Genbank No. U93215; the gene encoding oleosin from *Arabidopsis*, Genbank No. Z17657; Atmyc1 from *Arabidopsis*, Urao (1996) 20 *Plant Mol. Biol.* 32:571-576; the 2s seed storage protein gene family from *Arabidopsis*, Conceicao (1994) *Plant* 5:493-505; the gene encoding oleosin 20kD from *Brassica napus*, GenBank No. M63985; *napA* from *Brassica napus*, GenBank No. J02798, Josefsson (1987) *JBL* 26:12196-1301; the napin gene family from *Brassica napus*, Sjodahl (1995) *Planta* 197:264-271; the gene encoding the 2S storage protein from *Brassica napus*, Dasgupta (1993) *Gene* 133:301-302; the genes encoding oleosin a, Genbank No. U09118, 25 and, oleosin B, Genbank No. U09119, from soybean; and, the gene encoding low molecular weight sulphur rich protein from soybean, Choi (1995) *Mol Gen. Genet.* 246:266-268. The tissue specific E8 promoter from tomato is particularly useful for directing gene expression so that a desired gene product is located in fruits. Other suitable 30 promoters include those from genes encoding embryonic storage proteins.

One of skill will recognize that a tissue-specific promoter may drive expression of operably linked sequences in tissues other than the target tissue. Thus, as

used herein a tissue-specific promoter is one that drives expression preferentially in the target tissue, but may also lead to some expression in other tissues as well.

The invention also provides for use of tissue-specific promoters derived from viruses which can include, e.g., the tobamovirus subgenomic promoter (Kumagai 5 (1995) *Proc. Natl. Acad. Sci. USA* 92:1679-1683; the rice tungro bacilliform virus (RTBV), which replicates only in phloem cells in infected rice plants, with its promoter which drives strong phloem-specific reporter gene expression; the cassava vein mosaic virus (CVMV) promoter, with highest activity in vascular elements, in leaf mesophyll cells, and in root tips (Verdaguer (1996) *Plant Mol. Biol.* 31:1129-1139).

10 In some embodiments, the nucleic acid construct will comprise a promoter functional in a specific plant cell, such as in a species of *Lactuca*, operably linked to an RG polynucleotide. Promoters useful in these embodiments include RG promoters. In additional embodiments, the nucleic acid construct will comprise a RG promoter operably linked to a heterologous polynucleotide. The heterologous polynucleotide is chosen to 15 provide a plant with a desired phenotype. For example, the heterologous polynucleotide can be a structural gene which encodes a polypeptide which imparts a desired resistance phenotype. Alternatively, the heterologous polynucleotide may be a regulatory gene which might play a role in transcriptional and/or translational control to suppress, enhance, or otherwise modify the transcription and/or expression of an endogenous gene within the 20 plant. The heterologous polynucleotide of the nucleic acid construct of the present invention can be expressed in either sense or anti-sense orientation as desired. It will be appreciated that control of gene expression in either sense or anti-sense orientation can have a direct impact on the observable plant characteristics.

Modifying and Inhibiting RG Gene Expression

25 The invention also provides for RG nucleic acid sequences which are complementary to the RG polypeptide-encoding sequences of the invention; i.e., antisense RG nucleic acids. Antisense technology can be conveniently used to modify gene expression in plants. To accomplish this, a nucleic acid segment from the desired gene is cloned and operably linked to a promoter such that the anti-sense strand of RNA will be 30 transcribed. The construct is then transformed into plants and the antisense strand of RNA is produced. In plant cells, it has been shown that antisense RNA inhibits gene expression by preventing the accumulation of mRNA which encodes the enzyme of interest, see, e.g.,

Sheehy (1988) *Proc. Nat. Acad. Sci. USA* 85:8805-8809; Hiatt et al., U.S. Patent No. 4,801,340.

Antisense sequences are capable of inhibiting the transport, splicing or transcription of RG-encoding genes. The inhibition can be effected through the targeting of genomic DNA or messenger RNA. The transcription or function of targeted nucleic acid can be inhibited, e.g., by hybridization and/or cleavage. One particularly useful set of inhibitors provided by the present invention includes oligonucleotides which are able to either bind RG gene or message, in either case preventing or inhibiting the production or function of RG. The association can be though sequence specific hybridization. Such inhibitory nucleic acid sequences can, for example, be used to completely inhibit a plant disease resistance response. Another useful class of inhibitors includes oligonucleotides which cause inactivation or cleavage of RG message. The oligonucleotide can have enzyme activity which causes such cleavage, such as ribozymes. The oligonucleotide can be chemically modified or conjugated to an enzyme or composition capable of cleaving the complementary nucleic acid. One may screen a pool of many different such oligonucleotides for those with the desired activity.

Antisense Oligonucleotides

The invention provides for with antisense oligonucleotides capable of binding RG message which can inhibit RG activity by targeting mRNA. Strategies for designing antisense oligonucleotides are well described in the scientific and patent literature, and the skilled artisan can design such RG oligonucleotides using the novel reagents of the invention. In some situations, naturally occurring nucleic acids used as antisense oligonucleotides may need to be relatively long (18 to 40 nucleotides) and present at high concentrations. A wide variety of synthetic, non-naturally occurring nucleotide and nucleic acid analogues are known which can address this potential problem. For example, peptide nucleic acids (PNAs) containing non-ionic backbones, such as N-(2-aminoethyl) glycine units can be used. Antisense oligonucleotides having phosphorothioate linkages can also be used, as described in WO 97/03211; WO 96/39154; Mata (1997) *Toxicol Appl Pharmacol* 144:189-197; Antisense Therapeutics, ed. Agrawal (Humana Press, Totowa, N.J., 1996). Antisense oligonucleotides having synthetic DNA backbone analogues provided by the invention can also include phosphoro-dithioate, methylphosphonate,

phosphoramidate, alkyl phosphotriester, sulfamate, 3'-thioacetal, methylene(methylimino), 3'-N-carbamate, and morpholino carbamate nucleic acids, as described herein.

Combinatorial chemistry methodology can be used to create vast numbers of oligonucleotides that can be rapidly screened for specific oligonucleotides that have 5 appropriate binding affinities and specificities toward any target, such as the sense and antisense RG sequences of the invention (for general background information, see, e.g., Gold (1995) *J. of Biol. Chem.* 270:13581-13584).

Inhibitory Ribozymes

The invention provides for with ribozymes capable of binding RG message 10 which can inhibit RG activity by targeting mRNA. Strategies for designing ribozymes and selecting the RG-specific antisense sequence for targeting are well described in the scientific and patent literature, and the skilled artisan can design such RG ribozymes using the novel reagents of the invention. Ribozymes act by binding to a target RNA through the target RNA binding portion of a ribozyme which is held in close proximity to an enzymatic 15 portion of the RNA that cleaves the target RNA. Thus, the ribozyme recognizes and binds a target RNA through complementary base-pairing, and once bound to the correct site, acts enzymatically to cleave and inactivate the target RNA. Cleavage of a target RNA in such a manner will destroy its ability to direct synthesis of an encoded protein if the cleavage occurs in the coding sequence, or, preventing transport of the message from the nucleus to 20 the cytoplasm. After a ribozyme has bound and cleaved its RNA target, it is typically released from that RNA and so can bind and cleave new targets repeatedly.

Catalytic RNA molecules or ribozymes can also be used to inhibit 25 expression of any plant gene. It is possible to design ribozymes that specifically pair with virtually any target RNA and cleave the phosphodiester backbone at a specific location, thereby functionally inactivating the target RNA. In carrying out this cleavage, the ribozyme is not itself altered, and is thus capable of recycling and cleaving other molecules, making it a true enzyme. The inclusion of ribozyme sequences within antisense RNAs confers RNA-cleaving activity upon them, thereby increasing the activity of the constructs. The design and use of target RNA-specific ribozymes is described, e.g., in 30 Haseloff (1988) *Nature* 334:585-591.

In some circumstances, the enzymatic nature of a ribozyme can be advantageous over other technologies, such as antisense technology (where a nucleic acid

molecule simply binds to a nucleic acid target to block its transcription, translation or association with another molecule) as the effective concentration of ribozyme necessary to effect a therapeutic treatment can be lower than that of an antisense oligonucleotide. This potential advantage reflects the ability of the ribozyme to act enzymatically. Thus, a single
5 ribozyme molecule is able to cleave many molecules of target RNA. In addition, a ribozyme is typically a highly specific inhibitor, with the specificity of inhibition depending not only on the base pairing mechanism of binding, but also on the mechanism by which the molecule inhibits the expression of the RNA to which it binds. That is, the inhibition is caused by cleavage of the RNA target and so specificity is defined as the ratio
10 of the rate of cleavage of the targeted RNA over the rate of cleavage of non-targeted RNA. This cleavage mechanism is dependent upon factors additional to those involved in base pairing. Thus, the specificity of action of a ribozyme can be greater than that of antisense oligonucleotide binding the same RNA site.

The enzymatic ribozyme RNA molecule can be formed in a hammerhead
15 motif, but may also be formed in the motif of a hairpin, hepatitis delta virus, group I intron or RNaseP-like RNA (in association with an RNA guide sequence). Examples of such hammerhead motifs are described by Rossi (1992) *Aids Research and Human Retroviruses* 8:183; hairpin motifs by Hampel (1989) *Biochemistry* 28:4929, and Hampel (1990) *Nuc. Acids Res.* 18:299; the hepatitis delta virus motif by Perrotta (1992) *Biochemistry* 31:16;
20 the RNaseP motif by Guerrier-Takada (1983) *Cell* 35:849; and the group I intron by Cech U.S. Pat. No. 4,987,071. The recitation of these specific motifs is not intended to be limiting; those skilled in the art will recognize that an enzymatic RNA molecule of this invention has a specific substrate binding site complementary to one or more of the target gene RNA regions, and has nucleotide sequence within or surrounding that substrate
25 binding site which imparts an RNA cleaving activity to the molecule.

Sense Supression

Another method of suppression is sense suppression. Introduction of nucleic acid configured in the sense orientation has been shown to be an effective means by which to block the transcription of target genes. For an example of the use of this method
30 to modulate expression of endogenous genes see, Napoli et al., *The Plant Cell* 2:279-289 (1990), and U.S. Patent No. 5,034,323.

Cloning of RG Polypeptides

Synthesis and/or cloning of RG polynucleotides and isolated nucleic acid constructs of the present invention are provided by methods well known to those of ordinary skill in the art. Generally, the nomenclature and the laboratory procedures in recombinant DNA technology described below are those well known and commonly employed in the art. Standard techniques are used for cloning, DNA and RNA isolation, amplification and purification. Generally enzymatic reactions involving DNA ligase, DNA polymerase, restriction endonucleases and the like are performed according to the manufacturer's specifications. These techniques and various other techniques are generally performed according to Sambrook *et al.*, *Molecular Cloning - A Laboratory Manual*, Cold Spring Harbor Laboratory, Cold Spring Harbor, New York, (1989).

The isolation of RG genes may be accomplished by a number of techniques. For instance, oligonucleotide probes based on the sequences disclosed here can be used to identify the desired gene in a cDNA or genomic DNA library. To construct genomic libraries, large segments of genomic DNA are generated by random fragmentation, e.g. using restriction endonucleases, and are ligated with vector DNA to form concatemers that can be packaged into the appropriate vector. To prepare a cDNA library, mRNA is isolated from the desired organ, such as roots and a cDNA library which contains the *RG* gene transcript is prepared from the mRNA. Alternatively, cDNA may be prepared from mRNA extracted from other tissues in which RG genes or homologs are expressed.

The cDNA or genomic library can then be screened using a probe based upon the sequence of a cloned RG gene such as the genes disclosed herein. Probes may be used to hybridize with genomic DNA or cDNA sequences to isolate homologous genes in the same or different plant species.

Those of skill in the art will appreciate that various degrees of stringency of hybridization can be employed in the assay; and either the hybridization or the wash medium can be stringent. As the conditions for hybridization become more stringent, there must be a greater degree of complementarity between the probe and the target for duplex formation to occur. The degree of stringency can be controlled by temperature, ionic strength, pH and the presence of a partially denaturing solvent such as formamide. For example, the stringency of hybridization is conveniently varied by changing the polarity of the reactant solution through manipulation of the concentration of formamide within the range of 0% to 50%.

Alternatively, the RG nucleic acids of the invention can be amplified from nucleic acid samples using a variety of amplification techniques, such as polymerase chain reaction (PCR) technology, to amplify the sequences of the RG and related genes directly from genomic DNA, from cDNA, from genomic libraries or cDNA libraries. PCR and other *in vitro* amplification methods may also be useful, for example, to clone nucleic acid sequences that code for proteins to be expressed, to make nucleic acids to use as probes for detecting the presence of the desired mRNA in samples, for nucleic acid sequencing, or for other purposes.

Oligonucleotides can be used to identify and detect additional RG families and RG family species using a variety of hybridization techniques and conditions. Suitable amplification methods include, but are not limited to: polymerase chain reaction, PCR (PCR PROTOCOLS, A GUIDE TO METHODS AND APPLICATIONS, ed. Innis, Academic Press, N.Y. (1990) and PCR STRATEGIES (1995), ed. Innis, Academic Press, Inc., N.Y. (Innis)), ligase chain reaction (LCR) (Wu (1989) *Genomics* 4:560; Landegren (1988) *Science* 241:1077; Barringer (1990) *Gene* 89:117); transcription amplification (Kwoh (1989) *Proc. Natl. Acad. Sci. USA* 86:1173); and, self-sustained sequence replication (Guatelli (1990) *Proc. Natl. Acad. Sci. USA*, 87:1874); Q Beta replicase amplification and other RNA polymerase mediated techniques (e.g., NASBA, Cangene, Mississauga, Ontario); see Berger (1987) *Methods Enzymol.* 152:307-316, Sambrook, and Ausubel, as well as Mullis (1987) U.S. Patent Nos. 4,683,195 and 4,683,202; Arnheim (1990) *C&EN* 36-47; Lomell *J. Clin. Chem.*, 35:1826 (1989); Van Brunt, *Biotechnology*, 8:291-294 (1990); Wu (1989) *Gene* 4:560; Sooknanan (1995) *Biotechnology* 13:563-564. Methods for cloning *in vitro* amplified nucleic acids are described in Wallace, U.S. Pat. No. 5,426,039.

The degree of complementarity (sequence identity) required for detectable binding will vary in accordance with the stringency of the hybridization medium and/or wash medium. The degree of complementarity will optimally be 100 percent; however, it should be understood that minor sequence variations in the probes and primers may be compensated for by reducing the stringency of the hybridization and/or wash medium as described earlier.

In some preferred embodiments, members of this class of pest resistance genes can be identified by their ability to be amplified by PCR primers based on the sequences disclosed here. Appropriate primers and probes for identifying RG sequences

from plant tissues are generated from comparisons of the sequences provided herein. See, e.g., Table 1. For a general overview of PCR see *PCR Protocols: A Guide to Methods and Applications*. (Innis, M, Gelfand, D., Sninsky, J. and White, T., eds.), Academic Press, San Diego (1990), incorporated herein by reference.

5 Briefly, the first step of each cycle of the PCR involves the separation of the nucleic acid duplex formed by the primer extension. Once the strands are separated, the next step in PCR involves hybridizing the separated strands with primers that flank the target sequence. The primers are then extended to form complementary copies of the target strands. For successful PCR amplification, the primers are designed so that the
10 position at which each primer hybridizes along a duplex sequence is such that an extension product synthesized from one primer, when separated from the template (complement), serves as a template for the extension of the other primer. The cycle of denaturation, hybridization, and extension is repeated as many times as necessary to obtain the desired amount of amplified nucleic acid.

15 In the preferred embodiment of the PCR process, strand separation is achieved by heating the reaction to a sufficiently high temperature for an sufficient time to cause the denaturation of the duplex but not to cause an irreversible denaturation of the polymerase (see U.S. Patent No. 4,965,188). Template-dependent extension of primers in PCR is catalyzed by a polymerizing agent in the presence of adequate amounts of four
20 deoxyribonucleotide triphosphates (typically dATP, dGTP, dCTP, and dTTP) in a reaction medium comprised of the appropriate salts, metal cations, and pH buffering system. Suitable polymerizing agents are enzymes known to catalyze template-dependent DNA synthesis.

25 Polynucleotides may also be synthesized by well-known techniques as described in the technical literature. See, e.g., Carruthers *et al.*, *Cold Spring Harbor Symp. Quant. Biol.* 47:411-418 (1982), and Adams *et al.*, *J. Am. Chem. Soc.* 105:661 (1983). Double stranded DNA fragments may then be obtained either by synthesizing the complementary strand and annealing the strands together under appropriate conditions, or by adding the complementary strand using DNA polymerase with an appropriate primer
30 sequence.

The present invention further provides isolated RG proteins encoded by the RG polynucleotides disclosed herein. One of skill will recognize that the nucleic acid encoding a functional RG protein need not have a sequence identical to the exemplified genes disclosed here. For example, because of codon degeneracy a large number of 5 nucleic acid sequences can encode the same polypeptide. In addition, the polypeptides encoded by the RG genes, like other proteins, have different domains which perform different functions. Thus, the RG gene sequences need not be full length, so long as the desired functional domain of the protein is expressed.

The resistance proteins are at least 25 amino acid residues in length. 10 Typically, the RG proteins are at least 50 amino acid residues, generally at least 100, preferably at least 150, more preferably at least 200 amino acids in length. In particularly preferred embodiments, the RG proteins are of sufficient length to provide resistance to pests when expressed in the desired plants. Generally then, the RG proteins will be the length encoded by an RG gene of the present invention. However, those of ordinary skill 15 will appreciate that minor deletions, substitutions, or additions to an RG protein will typically yield a protein with pest resistance characteristics similar or identical to that of the full length sequence. Thus, full-length RG proteins modified by 1, 2, 3, 4, or 5 deletions, substitutions, or additions, generally provide an effective degree of pest resistance relative to the full-length protein.

20 The RG proteins which provide pest resistance will typically comprise at least one of an LRR or an NBS. Preferably, both are present. LRR and/or NBS regions present in the RG proteins of the present invention can be provided by RG genes of the present invention. In some embodiments, the LRR and/or NBS regions are obtained from other pest resistance genes. See, e.g., Yu *et al.*, *Proc. Natl. Acad. Sci. USA*, 93: 11751-25 11756 (1996); Bent *et al.*, *Science*, 265: 1856-1860 (1994).

Modified protein chains can also be readily designed utilizing various recombinant DNA techniques well known to those skilled in the art. For example, the chains can vary from the naturally occurring sequence at the primary structure level by amino acid substitutions, additions, deletions, and the like. Modification can also include 30 swapping domains from the proteins of the invention with related domains from other pest resistance genes.

Pests that can be targeted by RG genes and proteins of the present invention include such bacterial pests as *Erwinia carotovora* and *Pseudomonas marginalis*. Fungal pests which can be targeted by the present invention include *Bremia lactucae*, *Marssonina panattoniana*, *Rhizoctonia solani*, *Olpidium brassicae*, root aphid, *Sclerotinia sclerotiorum* and *S. minor*, and *Botrytis cinerea* which causes gray mold. RG genes also provide resistance to viral diseases such as lettuce and turnip mosaic viruses.

Fusion Proteins

RG polypeptides can also be expressed as recombinant proteins with one or more additional polypeptide domains linked thereto to facilitate protein detection, purification, or other applications. Such detection and purification facilitating domains include, but are not limited to, metal chelating peptides such as polyhistidine tracts and histidine-tryptophan modules that allow purification on immobilized metals, protein domains that allow purification on immobilized immunoglobulin, and the domain utilized in the FLAGS extension/affinity purification system (Immunex Corp, Seattle WA). The inclusion of a cleavable linker sequences such as Factor Xa or enterokinase (Invitrogen, San Diego CA) between the purification domain and plant disease resistant polypeptide may be useful to facilitate purification. One such expression vector provides for expression of a fusion protein comprising the sequence encoding a plant disease resistant polypeptide of the invention and nucleic acid sequence encoding six histidine residues followed by thioredoxin and an enterokinase cleavage site (e.g., see Williams (1995) *Biochemistry* 34:1787-1797). The histidine residues facilitate detection and purification while the enterokinase cleavage site provides a means for purifying the desired protein(s) from the remainder of the fusion protein. Technology pertaining to vectors encoding fusion proteins and application of fusion proteins are well described, see e.g., Kroll (1993) *DNA Cell. Biol.*, 12:441-53.

Antibodies Reactive to RG Polypeptides and Immunological Assays

The present invention also provides antibodies which specifically react with RG proteins of the present invention under immunologically reactive conditions. An antibody immunologically reactive with a particular antigen can be generated *in vivo* or by recombinant methods such as selection of libraries of recombinant antibodies in phage or similar vectors. "Immunologically reactive conditions" includes reference to conditions which allow an antibody, generated to a particular epitope of an antigen, to bind to that

epitope to a detectably greater degree than the antibody binds to substantially all other epitopes, generally at least two times above background binding, preferably at least five times above background. Immunologically reactive conditions are dependent upon the format of the antibody binding reaction and typically are those utilized in immunoassay
5 protocols.

"Antibody" includes reference to an immunoglobulin molecule obtained by *in vitro* or *in vivo* generation of the humoral response, and includes both polyclonal and monoclonal antibodies. The term also includes genetically engineered forms such as chimeric antibodies (e.g., humanized murine antibodies), heteroconjugate antibodies (e.g.,
10 bispecific antibodies), and recombinant single chain Fv fragments (scFv). The term "antibody" also includes antigen binding forms of antibodies (e.g., Fab', F(ab')₂, Fab, Fv, rIgG, and, inverted IgG). See, Pierce Catalog and Handbook, 1994-1995 (Pierce Chemical Co., Rockford, IL). An antibody immunologically reactive with a particular antigen can be generated *in vivo* or by recombinant methods such as selection of libraries
15 of recombinant antibodies in phage or similar vectors. See, e.g., Huse *et al.* (1989) *Science* 246:1275-1281; and Ward, *et al.* (1989) *Nature* 341:544-546; and Vaughan *et al.* (1996) *Nature Biotechnology*, 14:309-314.

Many methods of making antibodies are known to persons of skill. A number of immunogens are used to produce antibodies specifically reactive to an isolated
20 RG protein of the present invention under immunologically reactive conditions. An isolated recombinant, synthetic, or native RG protein of the present invention is the preferred immunogens (antigen) for the production of monoclonal or polyclonal antibodies.

The RG protein is then injected into an animal capable of producing antibodies. Either monoclonal or polyclonal antibodies can be generated for subsequent
25 use in immunoassays to measure the presence and quantity of the RG protein. Methods of producing monoclonal or polyclonal antibodies are known to those of skill in the art. See,
e.g., Coligan (1991) *Current Protocols in Immunology* Wiley/Greene, NY; and Harlow
and Lane (1989) *Antibodies: A Laboratory Manual* Cold Spring Harbor Press, NY);
Goding (1986) *Monoclonal Antibodies: Principles and Practice* (2d ed.) Academic Press,
30 New York, NY.

Frequently, the RG proteins and antibodies will be labeled by joining, either covalently or non-covalently, a substance which provides for a detectable signal. A wide

variety of labels and conjugation techniques are known and are reported extensively in both the scientific and patent literature. Suitable labels include radionucleotides, enzymes, substrates, cofactors, inhibitors, fluorescent moieties, chemiluminescent moieties, magnetic particles, and the like. Patents teaching the use of such labels include U.S. 5 Patent Nos. 3,817,837; 3,850,752; 3,939,350; 3,996,345; 4,277,437; 4,275,149; and 4,366,241.

The antibodies of the present invention can be used to screen plants for the expression of RG proteins of the present invention. The antibodies of this invention are also used for affinity chromatography in isolating RG protein.

10 The present invention further provides RG polypeptides that specifically bind, under immunologically reactive conditions, to an antibody generated against a defined immunogen, such as an immunogen consisting of the RG polypeptides of the present invention. Immunogens will generally be at least 10 contiguous amino acids from an RG polypeptide of the present invention. Optionally, immunogens can be from regions 15 exclusive of the NBS and/or LRR regions of the RG polypeptides. Nucleic acids which encode such cross-reactive RG polypeptides are also provided by the present invention. The RG polypeptides can be isolated from any number plants as discussed earlier. Preferred are species from the family *Compositae* and in particular the genus *Lactuca* such as *L. sativa* and such subspecies as *crispa*, *longifolia*, and *asparagina*.

20 "Specifically binds" includes reference to the preferential association of a ligand, in whole or part, with a particular target molecule (i.e., "binding partner" or "binding moiety") relative to compositions lacking that target molecule. It is, of course, recognized that a certain degree of non-specific interaction may occur between a ligand and a non-target molecule. Nevertheless, specific binding, may be distinguished as mediated 25 through specific recognition of the target molecule. Typically specific binding results in a much stronger association between the ligand and the target molecule than between the ligand and non-target molecule. Specific binding by an antibody to a protein under such conditions requires an antibody that is selected for its specificity for a particular protein. The affinity constant of the antibody binding site for its cognate monovalent antigen is at 30 least 10^7 , usually at least 10^8 , preferably at least 10^9 , more preferably at least 10^{10} , and most preferably at least 10^{11} liters/mole. A variety of immunoassay formats are appropriate for selecting antibodies specifically reactive with a particular protein. For

example, solid-phase ELISA immunoassays are routinely used to select monoclonal antibodies specifically reactive with a protein. See Harlow and Lane (1988) *Antibodies, A Laboratory Manual*, Cold Spring Harbor Publications, New York, for a description of immunoassay formats and conditions that can be used to determine specific reactivity. The antibody may be polyclonal but preferably is monoclonal. Generally, antibodies cross-reactive to such proteins as RPS2, RPM1 (bacterial resistances in *Arabidopsis*, L6 (fungal resistance in flax, PRF (resistance to *Pseudomonas syringae* in tomato), and N, (virus resistance in tobacco), are removed by immunoabsorption.

Immunoassays in the competitive binding format are typically used for cross-reactivity determinations. For example, an immunogenic RG polypeptide is immobilized to a solid support. Polypeptides added to the assay compete with the binding of the antisera to the immobilized antigen. The ability of the above polypeptides to compete with the binding of the antisera to the immobilized RG polypeptide is compared to the immunogenic RG polypeptide. The percent cross-reactivity for the above proteins is calculated, using standard calculations. Those antisera with less than 10% cross-reactivity with such proteins as RPS2, RPM1, L6, PRF, and N, are selected and pooled. The cross-reacting antibodies are then removed from the pooled antisera by immunoabsorption with these non-RG resistance proteins.

The immunoabsorbed and pooled antisera are then used in a competitive binding immunoassay to compare a second "target" polypeptide to the immunogenic polypeptide. In order to make this comparison, the two polypeptides are each assayed at a wide range of concentrations and the amount of each polypeptide required to inhibit 50% of the binding of the antisera to the immobilized protein is determined using standard techniques. If the amount of the target polypeptide required is less than twice the amount of the immunogenic polypeptide that is required, then the target polypeptide is said to specifically bind to an antibody generated to the immunogenic protein. As a final determination of specificity, the pooled antisera is fully immunosorbed with the immunogenic polypeptide until no binding to the polypeptide used in the immunosorption is detectable. The fully immunosorbed antisera is then tested for reactivity with the test polypeptide. If no reactivity is observed, then the test polypeptide is specifically bound by the antisera elicited by the immunogenic protein.

Production of transgenic plants of the invention

Isolated nucleic acid constructs prepared as described herein can be introduced into plants according techniques known in the art. In some embodiments, the introduced nucleic acid is used to provide RG gene expression and therefore pest resistance in desired plants. In some embodiments, RG promoters are used to drive expression of desired heterologous genes in plants. Finally, in some embodiments, the constructs can be used to suppress expression of a target endogenous gene, including RG genes.

To use isolated RG sequences in the above techniques, recombinant DNA vectors suitable for transformation of plant cells are prepared. Techniques for transforming a wide variety of higher plant species are well known and described in the technical and scientific literature. See, for example, Weising *et al. Ann. Rev. Genet.* 22:421-477 (1988).

A DNA sequence coding for the desired RG polypeptide, for example a cDNA or a genomic sequence encoding a full length protein, will be used to construct a recombinant expression cassette which can be introduced into the desired plant. An expression cassette will typically comprise the RG polynucleotide operably linked to transcriptional and translational initiation regulatory sequences which will direct the transcription of the sequence from the RG gene in the intended tissues of the transformed plant.

Such DNA constructs may be introduced into the genome of the desired plant host by a variety of conventional techniques. For example, the DNA construct may be introduced directly into the genomic DNA of the plant cell using techniques such as electroporation, PEG poration, particle bombardment and microinjection of plant cell protoplasts or embryogenic callus, or the DNA constructs can be introduced directly to plant tissue using ballistic methods, such as DNA particle bombardment. Alternatively, the DNA constructs may be combined with suitable T-DNA flanking regions and introduced into a conventional *Agrobacterium tumefaciens* host vector. The virulence functions of the *Agrobacterium tumefaciens* host will direct the insertion of the construct and adjacent marker into the plant cell DNA when the cell is infected by the bacteria.

Transformation techniques are known in the art and well described in the scientific and patent literature. The introduction of DNA constructs using polyethylene glycol precipitation is described in Paszkowski *et al. Embo J.* 3:2717-2722 (1984).

Electroporation techniques are described in Fromm *et al.* *Proc. Natl. Acad. Sci. USA* 82:5824 (1985). Ballistic transformation techniques are described in Klein *et al.* *Nature* 327:70-73 (1987).

Agrobacterium tumefaciens-mediated transformation techniques are well described in the scientific literature. See, for example Horsch *et al.* *Science* 233:496-498 (1984), and Fraley *et al.* *Proc. Natl. Acad. Sci. USA* 80:4803 (1983). Although Agrobacterium is useful primarily in dicots, certain monocots can be transformed by Agrobacterium. For instance, Agrobacterium transformation of rice is described by Hiei *et al.*, *Plant J.* 6:271-282 (1994). A particularly preferred means of transforming lettuce is described in Michelmore *et al.*, *Plant Cell Reports*, 6:439-442 (1987).

Transformed plant cells which are derived by any of the above transformation techniques can be cultured to regenerate a whole plant which possesses the transformed genotype and thus the desired RG-controlled phenotype. Such regeneration techniques rely on manipulation of certain phytohormones in a tissue culture growth medium, typically relying on a biocide and/or herbicide marker which has been introduced together with the RG nucleotide sequences. Plant regeneration from cultured protoplasts is described in Evans *et al.*, *Protoplasts Isolation and Culture, Handbook of Plant Cell Culture*, pp. 124-176, Macmillan Publishing Company, New York, 1983; and Binding, *Regeneration of Plants, Plant Protoplasts*, pp. 21-73, CRC Press, Boca Raton, 1985. Regeneration can also be obtained from plant callus, explants, organs, or parts thereof. Such regeneration techniques are described generally in Klee *et al.* *Ann. Rev. of Plant Phys.* 38:467-486 (1987).

The methods of the present invention are particularly useful for incorporating the RG polynucleotides into transformed plants in ways and under circumstances which are not found naturally. In particular, the RG polypeptides may be expressed at times or in quantities which are not characteristic of natural plants.

One of skill will recognize that after the expression cassette is stably incorporated in transgenic plants and confirmed to be operable, it can be introduced into other plants by sexual crossing. Any of a number of standard breeding techniques can be used, depending upon the species to be crossed.

The present invention further provides methods for detecting RG resistance genes in a nucleic acid sample suspected of comprising an RG resistance gene. The means by which the RG resistance gene is detected is not a critical aspect of the invention. For example, RG resistance genes can be detected by the presence of amplicons using RG resistance gene specific primers. Additionally, RG resistance genes can be detected by assaying for specific hybridization of an RG polynucleotide to an RG resistance gene. In some embodiments, the RG resistance gene can be amplified prior to the step of contacting the nucleic acid sample with the RG polynucleotide.

In a typical detection method, the nucleic acid sample is contacted with an RG polynucleotide to form a hybridization complex. The hybridization complex may be detected directly (e.g., in Southern or northern blots), or indirectly (e.g., by subsequent primer extension during PCR amplification). The RG polynucleotide hybridizes under stringent conditions to an RG polynucleotide of the invention. Formation of the hybridization complex is directly or indirectly used to indicate the presence of the RG resistance gene in the nucleic acid sample.

Detection of the hybridization complex can be achieved using any number of well known methods. For example, the nucleic acid sample, or a portion thereof, may be assayed by hybridization formats including but not limited to, solution phase, solid phase, mixed phase, or *in situ* hybridization assays. Briefly, in solution (or liquid) phase hybridizations, both the target nucleic acid and the probe or primer are free to interact in the reaction mixture. In solid phase hybridization assays, probes or primers are typically linked to a solid support where they are available for hybridization with target nucleic acid in solution. In mixed phase, nucleic acid intermediates in solution hybridize to target nucleic acids in solution as well as to a nucleic acid linked to a solid support. In *in situ* hybridization, the target nucleic acid is liberated from its cellular surroundings in such as to be available for hybridization within the cell while preserving the cellular morphology for subsequent interpretation and analysis. The following articles provide an overview of the various hybridization assay formats: Singer *et al.*, *Biotechniques* 4(3):230-250 (1986); Haase *et al.*, *Methods in Virology*, Vol. VII, pp. 189-226 (1984); Wilkinson, "The theory and practice of *in situ* hybridization" In: *In situ Hybridization*, Ed. D.G. Wilkinson. IRL Press, Oxford University Press, Oxford; and *Nucleic Acid Hybridization: A Practical Approach*, Ed. Hames, B.D. and Higgins, S.J., IRL Press (1987).

The effect of the modification of RG gene expression can be measured by detection of increases or decreases in mRNA levels using, for instance, Northern blots. In addition, the phenotypic effects of gene expression can be detected by measuring nematode, fungal, bacterial, viral, or other pest resistance in plants. Suitable assays for determining pest resistance are well known. Michelmore and Crute, *Trans. Br. mycol. Soc.*, 79(3): 542-546 (1982).

The means by which hybridization complexes are detected is not a critical aspect of the present invention and can be accomplished by any number of methods currently known or later developed. RG polynucleotides can be labeled by any one of several methods typically used to detect the presence of hybridized nucleic acids. One common method of detection is the use of autoradiography using probes labeled with ³H, ¹²⁵I, ³⁵S, ¹⁴C, or ³²P, or the like. The choice of radioactive isotope depends on research preferences due to ease of synthesis, stability, and half lives of the selected isotopes. Other labels include ligands which bind to antibodies labeled with fluorophores, chemiluminescent agents, and enzymes. Alternatively, probes can be conjugated directly with labels such as fluorophores, chemiluminescent agents or enzymes. The choice of label depends on sensitivity required, ease of conjugation with the probe, stability requirements, and available instrumentation. Labeling the RG polynucleotide is readily achieved such as by the use of labeled PCR primers.

The choice of label dictates the manner in which the label is bound to the probe. Radioactive probes are typically made using commercially available nucleotides containing the desired radioactive isotope. The radioactive nucleotides can be incorporated into probes, for example, by using DNA synthesizers, by nick translation with DNA polymerase I, by tailing radioactive DNA bases to the 3' end of probes with terminal deoxynucleotidyl transferase, by treating single-stranded M13 plasmids having specific inserts with the Klenow fragment of DNA polymerase in the presence of radioactive deoxynucleotides, dNTP, by transcribing from RNA templates using reverse transcriptase in the presence of radioactive deoxynucleotides, dNTP, or by transcribing RNA from vectors containing specific RNA viral promoters (e.g., SP6 promoter) using the corresponding RNA polymerase (e.g., SP6 RNA polymerase) in the presence of radioactive ribonucleotides rNTP.

The probes can be labeled using radioactive nucleotides in which the isotope resides as a part of the nucleotide molecule, or in which the radioactive component is attached to the nucleotide via a terminal hydroxyl group that has been esterified to a radioactive component such as inorganic acids, *e.g.*, ^{32}P phosphate or ^{14}C organic acids, 5 or esterified to provide a linking group to the label. Base analogs having nucleophilic linking groups, such as primary amino groups, can also be linked to a label.

Non-radioactive probes are often labeled by indirect means. For example, a ligand molecule is covalently bound to the probe. The ligand then binds to an anti-ligand molecule which is either inherently detectable or covalently bound to a detectable signal 10 system, such as an enzyme, a fluorophore, or a chemiluminescent compound. Enzymes of interest as labels will primarily be hydrolases, such as phosphatases, esterases and glycosidases, or oxidoreductases, particularly peroxidases. Fluorescent compounds include fluorescein and its derivatives, rhodamine and its derivatives, dansyl, umbelliferone, etc. Chemiluminescers include luciferin, and 2,3-dihydropthalazinediones, *e.g.*, luminol. 15 Ligands and anti-ligands may be varied widely. Where a ligand has a natural anti-ligand, namely ligands such as biotin, thyroxine, and cortisol, it can be used in conjunction with its labeled, naturally occurring anti-ligands. Alternatively, any haptenic or antigenic compound can be used in combination with an antibody.

Probes can also be labeled by direct conjugation with a label. For example, 20 cloned DNA probes have been coupled directly to horseradish peroxidase or alkaline phosphatase, (Renz, M., and Kurz, K. (1984) A Colorimetric Method for DNA Hybridization. *Nucl. Acids Res.* 12: 3435-3444) and synthetic oligonucleotides have been coupled directly with alkaline phosphatase (Jablonski, E., *et al.* (1986) Preparation of Oligodeoxynucleotide-Alkaline Phosphatase Conjugates and Their Use as Hybridization 25 Probes. *Nuc. Acids. Res.* 14: 6115-6128; and Li P., *et al.* (1987) Enzyme-linked Synthetic Oligonucleotide probes: Non-Radioactive Detection of Enterotoxigenic *Escherichia Coli* in Faecal Specimens. *Nucl. Acids Res.* 15:5275-5287).

Definitions

30 Units, prefixes, and symbols can be denoted in their SI accepted form. Numeric ranges are inclusive of the numbers defining the range. Unless otherwise indicated, nucleic acids are written left to right in 5' to 3' orientation, respectively. The

headings provided herein are not limitations of the various aspects or embodiments of the invention which can be had by reference to the specification as a whole. Accordingly, the terms defined immediately below are more fully defined by reference to the specification as a whole.

5 As used herein, the term "plant" includes reference to whole plants, plant organs (e.g., leaves, stems, roots, etc.), seeds and plant cells and progeny of same. The class of plants which can be used in the methods of the invention is generally as broad as the class of higher plants amenable to transformation techniques, including both monocotyledonous and dicotyledonous plants.

10 As used herein, "pest" includes, but is not limited to, viruses, fungi, nematodes, insects, and bacteria.

As used herein, "heterologous" is a nucleic acid that originates from a foreign species, or, if from the same species, is substantially modified from its original form. For example, a promoter operably linked to a heterologous structural gene is from a 15 species different from that from which the structural gene was derived, or, if from the same species, one or both are substantially modified from their original form.

As used herein, "RG gene," alternatively referred to as "RLG gene," is a gene encoding resistance to plant pests, such as viruses, fungi, nematodes, insects, and bacteria, and which hybridizes under stringent conditions and/or has at least 60% sequence 20 identity at the deduced amino acid level to the exemplified sequences provided herein. RG genes encode "RG polypeptides," alternatively referred to as "RLG polypeptides," which can comprise LRR motifs and/or NBS motifs. The RG polypeptides encoded by RG genes have at least 55% or 60% sequence identity, typically at least 65% sequence identity, preferably at least 70% sequence identity, often at least 75% sequence identity, more 25 preferably at least 80% sequence identity, and most preferably at least 90% sequence identity at the deduced amino acid level relative to the exemplary RG sequences provided herein. The term "RG family" or "RG family genus" or "genus" includes reference to a group of RG polypeptide sequence species that have at least 60% amino acid sequence identity, and, the nucleic acids encoding these polypeptides. The individual species of a 30 genus, i.e., the members of a family, typically are genetically mapped to the same locus.

As used herein, "RG polynucleotide" includes reference to a contiguous sequence from an RG gene of at least 18, 20, 25, 30, 40, or 50 nucleotides in length, up to

at least about 100 or at least about 200 nucleotides in length. In some embodiments, the polynucleotide is preferably at least 100 nucleotides in length, more preferably at least 200 nucleotides in length, most preferably at least 500 nucleotides in length. Thus, RG polynucleotide may be a RG gene or a subsequence thereof.

As used herein, "isolated," when referring to a molecule or composition, such as, for example, an RG polypeptide or nucleic acid, means that the molecule or composition is separated from at least one other compound, such as a protein, other nucleic acids (*e.g.*, RNAs), or other contaminants with which it is associated *in vivo* or in its naturally occurring state. Thus, an RG polypeptide or nucleic acid is considered isolated when it has been isolated from any other component with which it is naturally associated, *e.g.*, cell membrane, as in a cell extract. An isolated composition can, however, also be substantially pure. An isolated composition can be in a homogeneous state and can be in a dry or an aqueous solution. Purity and homogeneity can be determined, for example, using analytical chemistry techniques such as polyacrylamide gel electrophoresis (SDS-PAGE) or high performance liquid chromatography (HPLC).

The term "nucleic acid" or "nucleic acid molecule" or "nucleic acid sequence" refers to a deoxyribonucleotide or ribonucleotide oligonucleotide in either single- or double-stranded form. The term encompasses nucleic acids, *i.e.*, oligonucleotides, containing known analogues of natural nucleotides which have similar or improved binding properties, for the purposes desired, as the reference nucleic acid. The term also includes nucleic acids which are metabolized in a manner similar to naturally occurring nucleotides or at rates that are improved thereover for the purposes desired. The term also encompasses nucleic-acid-like structures with synthetic backbones. DNA backbone analogues provided by the invention include phosphodiester, phosphorothioate, phosphorodithioate, methylphosphonate, phosphoramidate, alkyl phosphotriester, sulfamate, 3'-thioacetal, methylene(methylimino), 3'-N-carbamate, morpholino carbamate, and peptide nucleic acids (PNAs); see Oligonucleotides and Analogues, a Practical Approach, edited by F. Eckstein, IRL Press at Oxford University Press (1991); Antisense Strategies, Annals of the New York Academy of Sciences, Volume 600, Eds. Baserga and Denhardt (NYAS 1992); Milligan (1993) *J. Med. Chem.* 36:1923-1937; Antisense Research and Applications (1993, CRC Press). PNAs contain non-ionic backbones, such as N-(2-aminoethyl) glycine units. Phosphorothioate linkages are described in WO

97/03211; WO 96/39154; Mata (1997) *Toxicol Appl Pharmacol* 144:189-197. Other synthetic backbones encompasses by the term include methyl-phosphonate linkages or alternating methylphosphonate and phosphodiester linkages (Strauss-Soukup (1997) *Biochemistry* 36:8692-8698), and benzylphosphonate linkages (Samstag (1996) *Antisense Nucleic Acid Drug Dev* 6:153-156). The term nucleic acid is used interchangeably with gene, cDNA, mRNA, oligonucleotide primer, probe and amplification product. Unless otherwise indicated, a particular nucleic acid sequence includes the complementary sequence thereof.

The term "exogenous nucleic acid" refers to a nucleic acid that has been isolated, synthesized, cloned, ligated, excised in conjunction with another nucleic acid, in a manner that is not found in nature, and/or introduced into and/or expressed in a cell or cellular environment other than or at levels or forms different than the cell or cellular environment in which said nucleic acid or protein is found in nature. The term encompasses both nucleic acids originally obtained from a different organism or cell type than the cell type in which it is expressed, and also nucleic acids that are obtained from the same cell line as the cell line in which it is expressed. invention.

The term "recombinant," when used with reference to a cell, or to the nucleic acid, protein or vector refers to a material, or a material corresponding to the natural or native form of the material, that has been modified by the introduction of a new moiety or alteration of an existing moiety, or is identical thereto but produced or derived from synthetic materials. For example, recombinant cells express genes that are not found within the native (non-recombinant) form of the cell or express native genes that are otherwise expressed at a different level, typically, under-expressed or not expressed at all. The term "recombinant means" encompasses all means of expressing, *i.e.*, transcription or translation of, an isolated and/or cloned nucleic acid *in vitro* or *in vivo*. For example, the term "recombinant means" encompasses techniques where a recombinant nucleic acid, such as a cDNA encoding a protein, is inserted into an expression vector, the vector is introduced into a cell and the cell expresses the protein. "Recombinant means" also encompass the ligation of nucleic acids having coding or promoter sequences from different sources into one vector for expression of a fusion protein, constitutive expression of a protein, or inducible expression of a protein, such as the plant disease resistant, or RG. polypeptides of the invention.

The term "specifically hybridizes" refers to a nucleic acid that hybridizes, duplexes or binds to a particular target DNA or RNA sequence. The target sequences can be present in a preparation of total cellular DNA or RNA. Proper annealing conditions depend, for example, upon a nucleic acid's, such as a probe's length, base composition, and the number of mismatches and their position on the probe, and can be readily determined empirically providing the appropriate reagents are available. For discussions of nucleic acid probe design and annealing conditions, see, *e.g.*, Sambrook and Ausubel.

The terms "stringent hybridization," "stringent conditions," or "specific hybridization conditions" refers to conditions under which an oligonucleotide (when used, for example, as a probe or primer) will hybridize to its target subsequence, such as an RG nucleic acid in an expression vector of the invention but not to a non-RG sequence.

Stringent conditions are sequence-dependent. Thus, in one set of stringent conditions an oligonucleotide probe will hybridize to only one species of the genus of RG nucleic acids of the invention. In another set of stringent conditions (less stringent) an oligonucleotide probe will hybridize to all species of the invention's genus but not to non-RG nucleic acids. Longer sequences hybridize specifically at higher temperatures. Stringent conditions are selected to be about 5⁰C lower than the thermal melting point (T_m) for the specific sequence at a defined ionic strength and pH. The T_m is the temperature (under defined ionic strength, pH, and nucleic acid concentration) at which 50% of the probes complementary to the target sequence hybridize to the target sequence at equilibrium (if the target sequences are present in excess, at T_m , 50% of the probes are occupied at equilibrium). Typically, stringent conditions will be those in which the salt concentration is less than about 1.0 M sodium ion, *i.e.*, about 0.01 to 1.0 M sodium ion concentration (or other salts) at pH 7.0 to 8.3 and the temperature is at least about 30⁰C for short probes (*e.g.*, 10 to 50 nucleotides) and at least about 60⁰C for long probes (*e.g.*, greater than 50 nucleotides). Stringent conditions may also be achieved with the addition of destabilizing agents such as formamide. Often, high stringency wash conditions preceded by low stringency wash conditions to remove background probe signal. An example of medium stringency wash conditions for a duplex of, *e.g.*, more than 100 nucleotides, is 1x SSC at 45⁰C for 15 minutes (see Sambrook for a description of SSC buffer). An example low stringency wash for a duplex of, *e.g.*, more than 100 nucleotides, is 4-6x SSC at 40⁰C for 15 minutes. a signal to noise ratio of 2x (or higher) than that observed for an unrelated

probe in the particular hybridization assay indicates detection of a "specific hybridization." Nucleic acids which do not hybridize to each other under stringent conditions can still be substantially identical if the polypeptides which they encode are substantially identical. This can occur, *e.g.*, when a nucleic acid is created that encodes for conservative substitutions. Stringent hybridization and stringent hybridization wash conditions are different under different environmental parameters, such as for Southern and Northern hybridizations. An extensive guide to the hybridization of nucleic acids is found in, *e.g.*, Sambrook, Tijssen (1993) *supra*.

As used herein "operably linked" includes reference to a functional linkage between a promoter and a second sequence, wherein the promoter sequence initiates and mediates transcription of the DNA sequence corresponding to the second sequence. Generally, operably linked means that the nucleic acid sequences being linked are contiguous and, where necessary to join two protein coding regions, contiguous and in the same reading frame.

In the expression of transgenes one of skill will recognize that the inserted polynucleotide sequence need not be identical and may be "substantially identical" to a sequence of the gene from which it was derived. As explained herein, these variants are specifically covered by this term.

In the case where the inserted polynucleotide sequence is transcribed and translated to produce a functional RG polypeptide, one of skill will recognize that because of codon degeneracy, a number of polynucleotide sequences will encode the same polypeptide. These variants are specifically covered by the term "RG polynucleotide sequence". In addition, the term specifically includes those full length sequences substantially identical (determined as described herein) with an RG gene sequence which encode proteins that retain the function of the RG protein. Thus, in the case of RG genes disclosed here, the term includes variant polynucleotide sequences which have substantial identity with the sequences disclosed here and which encode proteins capable of conferring resistance to nematodes, bacteria, viruses, fungi, insects or other pests on a transgenic plant comprising the sequence.

Two polynucleotides or polypeptides are said to be "identical" if the sequence of nucleotides or amino acid residues, respectively, in the two sequences is the same when aligned for maximum correspondence, as described below. The term

"complementary to" is used herein to mean that the complementary sequence is identical to all or a specified contiguous portion of a reference polynucleotide sequence.

The terms "sequence identity," "sequence similarity" and "homology" refer to when two sequences, such as the nucleic acid and amino acid sequences or the 5 polypeptides of the invention, when optimally aligned, as with, for example, the programs PILEUP, BLAST, GAP, FASTA or BESTFIT (see discussion, *supra*). "Percentage amino acid/nucleic acid sequence identity" refers to a comparison of the sequences of two polypeptides/nucleic acids which, when optimally aligned, have approximately the designated percentage of the same amino acids/nucleic acids, respectively. For example, 10 "60% sequence identity" and "60% homology" refer to a comparison of the sequences of two RG nucleic acids or polypeptides which, when optimally aligned, have 60% identity. For example, in one embodiment, nucleic acids encoding RG polypeptides of the invention comprise a sequence with at least 50% nucleic acid sequence identity to SEQ ID NO:1. In other embodiments, the RG polypeptides of the invention are encoded by nucleic acids 15 comprising a sequence with at least 50% sequence identity to SEQ ID NO:1, or, are encoded by nucleic acids comprising SEQ ID NO:1, or, have at least 60% amino acid sequence identity to the polypeptide of SEQ ID NO:2.

"Percentage of sequence identity" is determined by comparing two optimally aligned sequences over a comparison window, wherein the portion of the polynucleotide 20 sequence in the comparison window may comprise additions or deletions (i.e., gaps) as compared to the reference sequence (which does not comprise additions or deletions) for optimal alignment of the two sequences. The percentage is calculated by determining the number of positions at which the identical nucleic acid base or amino acid residue occurs in both sequences to yield the number of matched positions, dividing the number of 25 matched positions by the total number of positions in the window of comparison and multiplying the result by 100 to yield the percentage of sequence identity.

The term "substantial identity" of polynucleotide sequences means that a 30 polynucleotide comprises a sequence that has at least 55% or 60% sequence identity, generally at least 65%, preferably at least 70%, often at least 75%, more preferably at least 80% and most preferably at least 90%, compared to a reference sequence using the programs described above (preferably BESTFIT) using standard parameters. One of skill will recognize that these values can be appropriately adjusted to determine corresponding

identity of proteins encoded by two nucleotide sequences by taking into account codon degeneracy, amino acid similarity, reading frame positioning and the like. Substantial identity of amino acid sequences for these purposes normally means sequence identity of at least 55% or 60%, preferably at least 70%, more preferably at least 80%, and most preferably at least 95%. Polypeptides having "sequence similarity" share sequences as noted above except that residue positions which are not identical may differ by conservative amino acid changes. Conservative amino acid substitutions refer to the interchangeability of residues having similar side chains. For example, a group of amino acids having aliphatic side chains is glycine, alanine, valine, leucine, and isoleucine; a group of amino acids having aliphatic-hydroxyl side chains is serine and threonine; a group of amino acids having amide-containing side chains is asparagine and glutamine; a group of amino acids having aromatic side chains is phenylalanine, tyrosine, and tryptophan; a group of amino acids having basic side chains is lysine, arginine, and histidine; and a group of amino acids having sulfur-containing side chains is cysteine and methionine.

Preferred conservative amino acids substitution groups are: valine-leucine-isoleucine, phenylalanine-tyrosine, lysine-arginine, alanine-valine, and asparagine-glutamine.

Another indication that nucleotide sequences are substantially identical is if two molecules hybridize to each other under appropriate conditions. Appropriate conditions can be high or low stringency and will be different in different circumstances. Generally, stringent conditions are selected to be about 5°C to about 20°C lower than the thermal melting point (Tm) for the specific sequence at a defined ionic strength and pH. The Tm is the temperature (under defined ionic strength and pH) at which 50% of the target sequence hybridizes to a perfectly matched probe. Typically, stringent wash conditions are those in which the salt concentration is about 0.02 molar at pH 7 and the temperature is at least about 50°C. However, nucleic acids which do not hybridize to each other under stringent conditions are still substantially identical if the polypeptides which they encode are substantially identical. This may occur, *e.g.*, when a copy of a nucleic acid is created using the maximum codon degeneracy permitted by the genetic code.

Nucleic acids of the invention can be identified from a cDNA or genomic library prepared according to standard procedures and the nucleic acids disclosed here used as a probe. Thus, for example, stringent hybridization conditions will typically include at least one low stringency wash using 0.3 molar salt (*e.g.*, 2X SSC) at 65°C. The washes

are preferably followed by one or more subsequent washes using 0.03 molar salt (e.g., 0.2X SSC) at 50°C, usually 60°C, or more usually 65°C. Nucleic acid probes used to identify the nucleic acids are preferably at least 100 nucleotides in length.

As used herein, "nucleotide binding site" or "nucleotide binding domain" ("NBS") includes reference to highly conserved nucleotide-, *i.e.*, ATP/GTP-, binding domains, typically included in the "kinase domain" of kinase polypeptides, such as a kinase-1a, kinase 2, or a kinase 3a motif, as described herein. For example, the tobacco N and *Arabidopsis* RPS2 genes, among several recently cloned disease-resistance genes, share highly conserved NBS sequence. Kinase NBS subdomains further consist of three subdomain motifs: the P-loop, kinase-2, and kinase-3a subdomains (Yu (1996) *Proc. Acad. Sci. USA* 93:11751-11756). As discussed in detail herein, examples include the *Arabidopsis* RPP5 gene (Parker (1997) *supra*), the *A. thaliana* RPS2 gene (Mindrinos (1997) *supra*), and the flax L6 rust resistance gene (Lawrence (1995) *supra*) which all encode proteins containing an NBS; and Mindrinos (1994) *Cell* 78:1089-1099; and Shen (1993) *FEBS* 335:380-385. Using the teachings disclosed and incorporated herein and standard nucleic acid hybridization and/or amplification techniques, one of skill can identify members having NBS domains, including any of the genus of NBS-containing plant disease resistant polypeptides of the invention.

As used herein, "leucine rich region" ("LRR") includes reference to a region that has a leucine content of at least 20% leucine or isoleucine, or 30% of the aliphatic residues: leucine, isoleucine, methionine, valine, and phenylalanine, and arranged with approximate repeated periodicity. The length of the repeat may vary in length but is generally about 20 to 30 amino acids. An LRR-containing polypeptide typically will have the canonical 24 amino acid leucine-rich repeat (LRR) sequence, which is present in different proteins that mediates molecular recognition and/or interaction processes; as described in Bent (1994) *Science* 265:1856-1860; Parker (1997) *Plant Cell*. 9:879-894; Hong (1997) *Plant Physiol.* 113:1203-1212; Schmitz (1997) *Nucleic Acids Res.* 25:756-763; Hipskind (1996) *Mol. Plant Microbe Interact.* 9:819-825; Tornero (1996) *Plant J.* 10:315-330; Dixon (1996) *Cell* 84:451-459; Jones (1994) *Science* 266:789-793; Lawrence (1995) *Plant Cell* 7:1195-1206; Song (1995) *Science* 270:1804-1806; as discussed in further detail *supra*. Using the teachings disclosed and incorporated herein and standard nucleic acid hybridization and/or amplification techniques, one of skill can

identify polypeptides having LRR domains, including any member of the genus of LRR-containing RG polypeptides of the invention.

The term "promoter" refers to a region or sequence determinants located upstream or downstream from the start of transcription and which are involved in 5 recognition and binding of RNA polymerase and other proteins to initiate transcription. A "plant promoter" is a promoter capable of initiating and/or regulating transcription in plant cells; see also discussion on plant promoters, *supra*.

The term "constitutive promoter" refers to a promoter that initiates and helps control transcription in all tissues. Promoters that drive expression continuously 10 under physiological conditions are referred to herein as "constitutive" promoters and are active under most environmental conditions and states of development or cell differentiation; see also detailed discussion, *supra*.

The term "inducible promoter" refers to a promoter which directs transcription under the influence of changing environmental conditions or developmental 15 conditions. Examples of environmental conditions that may effect transcription by inducible promoters include anaerobic conditions, elevated temperature, drought, or the presence of light. Such promoters are referred to herein as "inducible" promoters; see also detailed discussion, *supra*.

The term "abscission-induced promoter" or "abscission promoter" refers to a 20 class of promoters which are activated upon plant ripening, such as fruit ripening, and are especially useful incorporated in the expression systems (*e.g.*, expression cassettes, vectors) of the invention. When the plant disease resistant polypeptide-encoding nucleic acid is under the control of an abscission promoter, rapid cell death, induced by expression 25 of the invention's polypeptide, accelerates and/or accentuates abscission of the plant part, increasing the efficiency of the harvesting of fruits or other plant parts, such as cotton, and the like; see also detailed discussion, *supra*.

The term "tissue-specific promoter" refers to a class of transcriptional control elements that are only active in particular cells or tissues. Examples of plant 30 promoters under developmental control include promoters that initiate transcription only (or primarily only) in certain tissues, such as roots, leaves, fruit, ovules, seeds, pollen, pistils, or flowers; see also detailed discussion, *supra*.

As used herein "recombinant" includes reference to a cell, or nucleic acid, or vector, that has been modified by the introduction of a heterologous nucleic acid or the alteration of a native nucleic acid to a form not native to that cell, or that the cell is derived from a cell so modified. Thus, for example, recombinant cells express genes that are not found within the native (non-recombinant) form of the cell or express native genes that are otherwise abnormally expressed, under expressed or not expressed at all.

As used herein, a "recombinant expression cassette" or "expression cassette" is a nucleic acid construct, generated recombinantly or synthetically, with a series of specified nucleic acid elements which permit transcription of a particular nucleic acid in a target cell. The expression vector can be part of a plasmid, virus, or nucleic acid fragment. Typically, the recombinant expression cassette portion of the expression vector includes a nucleic acid to be transcribed, and a promoter.

As used herein, "transgenic plant" includes reference to a plant modified by introduction of a heterologous polynucleotide. Generally, the heterologous polynucleotide is an RG structural or regulatory gene or subsequences thereof.

As used herein, "hybridization complex" includes reference to a duplex nucleic acid sequence formed by selective hybridization of two single-stranded nucleic acids with each other.

As used herein, "amplified" includes reference to an increase in the molarity of a specified sequence. Amplification methods include the polymerase chain reaction (PCR), the ligase chain reaction (LCR), the transcription-based amplification system (TAS), the self-sustained sequence replication system (SSR). A wide variety of cloning methods, host cells, and *in vitro* amplification methodologies are well-known to persons of skill.

As used herein, "nucleic acid sample" includes reference to a specimen suspected of comprising RG resistance genes. Such specimens are generally derived, directly or indirectly, from lettuce tissue.

The term "antibody" refers to a polypeptide substantially encoded by an immunoglobulin gene or immunoglobulin genes, or fragments or synthetic or recombinant analogues thereof which specifically bind and recognize analytes and antigens, such as a genus or subgenus of polypeptides of the invention, as described *supra*.

It is understood that the examples and embodiments described herein are for illustrative purposes only and that various modifications or changes in light thereof will be suggested to persons skilled in the art and are to be included within the spirit and purview of this application and scope of the appended claims.

5

EXAMPLES

The following examples are offered to illustrate, but not to limit the claimed invention.

Example 1

10 Example 1 describes the use of PCR to amplify RG genes from lettuce.

Multiple primers with low degeneracy, particularly at the 3' end, were designed based on the sequences of two known resistance genes from tobacco and flax.

DNA Templates

15 Lettuce genomic DNA was extracted from cultivar Diana and a mutant line derived from cultivar Diana using a standard CTAB protocol. To generate cDNA templates, RNA was isolated from cultivar Diana and the mutant following standard procedures; first strand cDNA was synthesized using Superscript reverse transcriptase from 1 Φ g total RNA as specified by the manufacturer (Life Technologies). BAC (bacterial artificial chromosome) clones from the *Dm3* region were isolated from a BAC library of 20 over 53,000 clones using marker AC15 that was known to be closely linked to *Dm3*. Bacterial plasmids containing clones of *L6* and *RPS2* were used as positive controls.

PCR with degenerate oligonucleotide primers

25 Oligonucleotide primers were designed based on conserved motifs in the nucleotide binding sites (NBS) of *L6*, *RPS2*, and *N*. Eight primers were made corresponding to the GVGKTT motif in the sense direction; each had 64-fold degeneracy. Six primers were made to the GPLAL motif in the anti-sense direction; with either 16 or 256-fold degeneracy (Table 1).

30 Oligonucleotides included 14-mer adaptors of (CUA)₄ at the 5' end of the sense primers and (CAU)₄ at the 5' end of the antisense primers to allow rapid cloning of the PCR products into pAMP1 (Life Technologies).

PCR amplification was performed in 50 μ l reaction volume with 1 μ M of each of a pair of sense and antisense primers. The templates were denatured by heating to 94EC for 2 min. This was followed by 35 cycles of 30 sec at 94EC, 1 min at 50EC, 2 min at 72EC, with a single final extension of 5 min at 72EC. 25 ng of genomic DNA or cDNA was used. BAC clones as templates required less. The final dNTP concentration was 0.2 mM; MgCl₂ was 1.5 mM.

Forty-eight combinations of sense and antisense primers were tested on a panel of nine templates consisting of two genomic DNA samples, two cDNA preparations, three BAC clones and plasmids containing *L6* and *RPS2* as positive controls.

10 Amplification from *L6* and *RPS2* resulted in fragments of 516 and 513 respectively. Seven combinations of primers resulted in fragments of approximately this size with multiple templates (Table 2). Primers that gave RLG products were: PLOOPAA, PLOOPAG, PLOOPGA, PLOOPGG, PLOOPAC, GLPL3, GLPL4.

15

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Table 1

DEGENERATE PRIMER SEQUENCES for NBS PCR

Sense primers based on GVGKTT amino acid sequence from L6, N and rps2 PLOOP motif:

PLOOPAG 5' GGN GTN GGN AAA ACG AC 3'

PLOOPAA 5' GGN GTN GGN AAA ACA AC 3'

PLOOPAT 5' GGN GTN GGN AAA ACT AC 3'

PLOOPAC 5' GGN GTN GGN AAA ACC AC 3'

PLOOPGG 5' GGN GTN GGN AAG ACG AC 3'

PLOOPGA 5' GGN GTN GGN AAG ACA AC 3'

PLOOPGT 5' GGN GTN GGN AAG ACT AC 3'

PLOOPGC 5' GGN GTN GGN AAG ACC AC 3'

Antisense primers based on GLPLAL amino acid sequence:

GLPL1 5' AGN GCN AGN GGN AGG CC 3'

GLPL2 5' AGN GCN AGN GGN AGA CC 3'

GLPL3 5' AGN GCN AGN GGN AGT CC 3'

GLPL4 5' AGN GCN AGN GGN AGC CC 3'

GLPL5 5' AAN GCC AAN GGC AAA CC 3'

GLPL6 5' AAN GCC AAN GGC AAT CC 3'

TABLE 2. Characteristics of RLGs isolated from lettuce.

	Template	Primers	Number ^a	Size ^b (bp)	Copy number ^c	Dm linkage
5	RLG1	genomic DNA	PLOOPGA+GLPL6	6/6	522	<i>DM4,</i> <i>DM13</i>
		cDNA	PLOOPGA+GLPL6	1/5		
		genomic DNA	PLOOPAA+GLPL6	5/5		
		cDNA	PLOOPAA+GLPL6	1/1		
10	RLG2	BACH8	PLOOPGG+GLPL3	3/3	510	<i>DM1,</i> <i>Dm3</i>
	RLG3	gemonic DNA	PLOOPGA+GLPL4	3/6	461	
	RLG4	genomic DNA	PLOOPGA+GLPL4	1/6	524	

^a Number of RLG sequences out of total number of clones sequenced.

^b Size of fragment amplified from the nucleotide bindind domain.

^c Estimated copy number from genomic Southern blot analysis and numbers of clones in the BAC library.

Example 2

Example 2 describes the genetic analysis used to obtain a preliminary indication of the linkage relationships of the amplified products and known clusters of resistance genes.

Bulked segregant analysis was performed to obtain a preliminary indication of the linkage relationships of the amplified products and known clusters of resistance genes. DNA from individuals were pooled for each susceptible and resistant bulk. Amplified products were then mapped by RFLP analysis from our intraspecific mapping population. Resistances from four clusters of resistance genes as well as over six hundred markers have now been mapped on this population. Linkage analysis was done using JIONMAP or MAPMAKER mapping programs. Due to a suppression of recombination in the *Dm3* region, sequences were mapped relative to *Dm3* using a panel of deletion mutants that provided greater genetic resolution than the mapping population (Anderson *et al.* 1996). All blots were washed twice at 63EC in 2x SSC/1% SDS for 20 min, followed by one wash at 63EC in 1x SSC/0.1% SDS for 10 or 30 min.

Most of the RLG sequences were analyzed by bulked segregant analysis (BSA) using pools of resistant and susceptible individuals for each of the four clusters of resistance genes. In genomic Southern analyses, all the RLGs revealed numerous fragments of varying intensity. The numbers of bands was highly dependent of the 5 stringency of hybridization. BSA demonstrated that RLG1 was linked to the *Dm4,7* and *Dm1,3* clusters. Segregation analysis confirmed this linkage.

RLG2 was derived from BAC H8 that was known to be from the *Dm3* region. BSA with RLG2 demonstrated that the polymorphic bands that distinguished the parents of our mapping population mapped to the *Dm1,Dm3* cluster. Several bands 10 absolutely cosegregated with *Dm1* or *Dm3*. To provide finer genetic resolution, RLG2 was also mapped using a panel of *Dm3* deletion mutants. A number of fragments were missing in largest deletion mutant demonstrating that several RLG2 family members are physically located very close to *Dm3*. No fragment was missing in all deletion mutants; however, this is not unexpected as there is extensive duplication within the region.

15

Example 3

Example 3 describes the screening of a bacterial artificial chromosome library.

Over 53,000 BAC clones containing lettuce genomic DNA were screened 20 with two of the amplified products. High density filters each containing 1536 clones were hybridized to ^{32}P labelled probes. Filters were washed at 65EC with 40 mM Na_2PO_4 /0.1% SDS for 5 min followed by 20 min in the same solution.

To isolate additional RLG sequences we screened our genomic BAC library. Clones were identified that hybridized to RLG1 and RLG2. Nearly all the clones that 25 hybridized to RLG2 also hybridized to marker AC15 that had already been shown by deletion mutant analysis to be clustered around *Dm3*. This provided further evidence for clustering of RLG2 sequences.

Using primers conserved within each family, part of the NBS was amplified 30 from each unique BAC clone and sequenced. This revealed that members within each family varied from 64% identical at the deduced amino acid level. The most divergent members only weakly cross-hybridized to each other. Currently, RLG sequences are

considered to be part of the same family of sequences if they are at least 55% identical at the deduced amino acid level and map to the same region of the chromosome.

Example 4:

5 Example 4 describes the cloning, identification, sequencing and characterization of RG polynucleotide sequences; including use of RG sequences from plasmid and PCR products.

Doubled stranded plasmid DNA clones and PCR products were sequenced using an ABI377 automated sequencer and fluorescently labelled di-deoxy terminators.
10 Sequences were assembled using Sequencher (Genecodes), DNASTar (DNASTar) and Genetics Computer Group (GCG, Madison, WI) software. Database searches were performed using BLASTX and FASTA (GCG) algorithms.

Sequences flanking the NBS region for RLG2 and for some of RLG1 were obtained by a series of IPCR and the products sequenced directly. IPCR worked less well
15 for RLG1. Therefore RLG1 was subcloned from a BAC clone into pBSK (Stratagene) and the double stranded plasmid sequenced by long range sequencing.

Initially, a total of 30 clones were sequenced. Three of these seven primer combinations yielded sequences that comprised continuous open reading frames with sequence identity to the NBS of known resistance genes. Seven out of 10 clones amplified
20 from genomic DNA with the primer pair PLOOPGA/GLP6 were 522 bp long; they were identical to each other and named RLG1. All six clones amplified from genomic DNA or cDNA using the primers PLOOPAA/GLP6 were similar/the same as RLG1. All three clones sequenced from BAC clone H8 were 510 bp long, identical to each other but different from RLG1 and were therefore designated RLG2. The 11 clones sequenced from
25 four other primer combinations had no similarity to any NBS motifs and therefore were not studied further. Therefore, sequencing resulted in the identification of clones containing NBS motifs representing four RLG sequences.

Comparison of the deduced amino acid sequences of RLG1 and RLG2 to those of known resistance genes revealed that RLG1 and RLG2 are as similar to each other
30 as they are to resistance genes from other species and that this is the same level of identity shown between the known resistance genes (Table 3). The percent identity (upper quadrant) and percent identity (lower quadrant) were determined using the MEGALIGN

routine of the DNASTAR package. Identity refers to the proportion of identical amino acids; identity refers to the proportion of identical and similar amino acids and takes into account substitutions of amino acids with similar chemical characteristics. RG1 and RG2 are as similar to each other and to cloned resistance genes as cloned resistance genes from 5 a variety of species are to each other. L6, resistance to *Melampsora lini* in flax (Lawrence *et al.*, 1995). N, resistance to tobacco mosaic virus in tobacco (Whitham *et al.*, 1994). PRF, required for resistance to *Pseudomonas syringae* in tomato. RPS2, resistance to *Pseudomonas syringae* in *Arabidopsis thaliana* (Bent *et al.*, 1994; Mindrinos *et al.*, 1994). RPM1, resistance to *Pseudomonas syringae* pv. *maculicola* in *A. thaliana* (Grant *et al.*, 10 1995). The initial RG1 and RG2, sequences were amplified from lettuce using degenerate primers.

15 **Table 3**
IDENTITIES OF
RESISTANCE GENE HOMOLOGUES

	RG1	RG2	RG3	RG4	N gene	RPS2
Lettuce	RG1	***	22.7	15.0	29.2	25.4
Lettuce	RG2		***	32.2	21.6	22.7
Lettuce	RG3			***	17.2	15.0
20 Lettuce	RG4				***	32.8
Tobacco	N gene					22.7
<i>Arabidopsis</i>	RPS2					21.6

25 The regions homologous to the primers are included in this analysis as the genomic sequences for RLG1 and RLG2 were determined by IPCR. Interestingly, the genomic sequences for RLG1 exactly matched that of the primers used.

To obtain further evidence that we had amplified resistance genes, we 30 amplified the regions flanking the NBSs of RLG1a and RLG2a by IPCR of BAC clones. These products were then directly sequenced without cloning to minimize the introduction of PCR artifacts. Sequence analysis of the 5' regions failed to detect any homology to known resistance genes. However, the sequence of the 3' region contained leucine-rich

repeats (LRRs). When this sequence was used to search GENBANK using BLASTX, it detected identity to the *Arabidopsis* resistance gene, *RPS2*. This region does not contain as regular LRRs as in some resistance genes; however, the repeat structure seems to be consistent with that of the flax resistance gene, *L6*. Therefore, the presence of an LRR 5 region is further evidence that the sequences we amplified using degenerate oligonucleotide primers are probably resistance genes.

The sequences of the IPCR products also provided the genomic sequences of the regions complementary to the sequences of the degenerate oligonucleotide primers. The genomic sequences for RLG1 were identical to one of the primers in the mixture. 10 The RLG sequences are resistance genes as supported by three criteria: the presence of multiple sequence motifs characteristic of resistance genes, genetic cosegregation with known resistance genes, and their existence as clustered multi-gene families. The presence of LRR regions in a similar position relative to the NBS as in cloned resistance genes provides stronger evidence than relying solely sequence similarity between NBS regions. 15 The clustering of RLG sequences at the same position as the known clusters of resistance genes make them strong candidates for encoding resistance genes. The hybridization patterns and genetic distribution of the RLG sequences are similar to that of cloned resistance genes in other species. Most of these hybridize to small multigene families and preliminary genetic evidence indicates that they are clustered in the genome. Therefore, 20 the degenerate primers that we designed from other resistance genes seemed to have been specific enough to amplify resistance genes rather than P-loop containing proteins in general.

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SEQ ID No: 1

RLG1A
[Strand]

1 ATCGTAACCGTTCTACGAG ANCGCTGTCCCCCTTCATC TTTTGTCTATACTCATATTC TCATNNAATTMTGCCACATNT
 81 AATTTTGTTTATTTAA TTAATTTTTATTCACATGT CATTTTATGAGTTTTCTAT TTATGAGTTCACTATAAT
 161 ATTTCATGTAATACAATA AATGCAATTATTTCTT TAAATAAACGCTATAATAT ATAGATTAATTCATATAAT
 241 ACATAGTTAAACTCATATA ATACATATGTCATCCCCAG TTATTTATATGTCATCC TTATTTATTTATTTAT
 321 TTATTAAGTAGATGATCTT TGTGATATAAAAATTAAAT TTGTTCAAATTTAAATTAAT TTAAATAATCCCACATTG
 401 ATAAAATTTAAAAAAATGGN CCCACCATTCGCTTCACT TTTTCACTCATCAATATCC TGAGTATTCCTTCGTTTC
 481 CACCCCTATCAATTTCGA CGAAATGACAGACTCTTCAG GCGTTCTGAATTGGCTTC CGACACTGTTCACTGAAGGA
 561 GATAATAATCAATGGAGC TGCTCCATGTCATTGCTG ATGAAAGGTGAATTTGATGT GAAGANAATGTCAGCGATCN
 641 ATCTCCATCGGAACCCACC ACATTATCAGTGTACCCCA ACCACTCRAAACGGYGGAA GTAGRRAKACWRKAAAGTCA
 721 TGAAGAATAGATTATTTTG TCTTCATGGCTGACTGAGG AGCGGGTTAGTTCATCATT TTCTTGANCAAAAGAATTA
 801 TCGGTCTACGAAATTTCAC ATCGACAAAGAAGTTTCAT TCGCAATGTTTGTAAACA ATTTTAACTTTTATCTT
 881 TTCATTGAAACTCTCAATT GCAATTGCAACTTCAACT TTGGGCCACAAATTGTTG TGCGCGTTAATTAAATCCA
 961 CATATTCACTGAAACAATA ATTCAAAATGATCTGTTG ATCCAATTCTACACATCTC TTGATAATTGAAATCATTCA
 1041 CGCTTCATCCATTTCATCCA CATCTATACATATTCTCTG CTCTTATCATATTAAACGAT GGCCTGAAATCTCTTCCTG
 1121 CCTTCTTGACAGTGGTTT GAAAGCTGGCATYTGAAGC TTGGAAGAAGTGTGCTC CAAAAGAATGAACTCTGAG
 1201 CTTAAGAAATTGAAAGGAGAC ATTGACCAAATCCAGATC TCTTAAAGATGCTTCCAG AAGGAAGTAACCTAATGAA
 1281 CGTTAAAGATGGCTGAATG ATCTCCAACTTGGCTTAT GACATAGACGGACTCTCTGA TGATTTGCAACTGAA
 1361 TTCAWCCTGAGTTGACCGAG GAGGGTGGAGCCTCCCTCCAG TATGGTAAGAAAACAACTATCC CAAGTTGTTGCAAGTTTC
 1441 TCACAAAGTAATAGGATGCA TGCCCAAGTTAGATGATATTG CCACCAAGTTTACAAGAATG GTAGAGGCAAAAATAATG
 1521 TGGTTTAAGTGTGATAACAT ATGAAAAGCCAAAATTGGA AGGTATGAGGGCTCTTGGT AGATGAAAGGGTACTGTG
 1601 GAGCTGAAGTAGTAAAGAAA AAATTCTGAGAAGCTGTG GGGGATAAAGTGAATCTAG GGAGTCAAAATCTAGCATC
 1681 TGCCCCATAGTTGGTATGGG TTGGAGTTGTTAAACACTC TAGCTAGACTTTGATGAT GAAAAGAAAGTGAAGGATCA
 1761 CTTCGAACCTCAGGGCTTGGG TTGGTGTCTGATGAGTTG AGTGTCCCAATATAAGCAG AGTTATTATCAATCTGTG
 1841 CTGGGGAAAAGGAGGAGTTT GAAGACTTAAATCTGCTTCAGA AGAAGCTCTTAAAGAGAAC TTAGGAACCAAGCTTCTA
 1921 ATAGTTTGGATGATGTTG GTCTGAAAGCTATGGTGTGTTT RTRTTCTGATAYTGTCTC TGTTTCCCAAGGACTATGAG
 2001 AAGTAGAATATCATGACAA CTGGCAAGGAGCAATTGCTC AGAAAGCTGGCTTCTCA TCAAGACCCCTCTGAGGGTC
 2081 TAATCACAAAGATGATGCTTGT TCTTGTGTTGCTCAACACGG ATTGGTGTACCAAACCTTG ATTCACTCCAACTAAGG
 2161 CCACATGGAGAAGCTTGTGTA GAAGAAATGATGGCTTAC CTCTAGCTTAAAGAACACTT GGAAGGTTTAAAGGACAAA
 2241 AACAGACGAGGACAAATGGA AGGAGCTGTTGAGTGTG ATATGGAGGTTAGGAAGAG CGATGAGATTGTCGGGCTC
 2321 TTGACTAAGCTACATGAT CTTCTGCTCCTTGAAGCT RTTRTTCTGATAYTGTCTC TGTTTCCCAAGGACTATGAG
 2401 TTGCAAGGAGGAGTTGAT TCTATTGTTGAGTGGCAGAAG GGGTTTGCAACCAACT AYAAACAAGTCACAGAACG
 2481 TTGGGTCTGAAATTTCATR AAGAGTTTGTCAAGRTCR TTTTTCAACATGCTCTAA TRRCAAATCSTGTTGTA
 2561 TGCATGACCTATGAAATGAT TTGGCTACATTGTTGCTCG AGAATTTTTTCAAGGTTAG ACATAGAGATGAAAGGAA
 2641 TTAGGATGSAATCTTGGG RAAGCACCGMCATATGTCAT TTGATGAGGTTAGTACATA GGTTACAAAARGTTGAGCC
 2721 ATTAGGAGGAGCTAAAATT TGAGAACATTTTAGCATG TCTGTTGGGGTGGTAGAAGA TTGGAAGATTTTACTTAT
 2801 CAAACAAGGTCTGAAATGAC WTACTTCARGAATTACCAT TTTAAGGGTCTRACTTTGA TTRRTCTTAYAATAASYRAG
 2881 GTACCARAACTCGTSGGTAG TATGAAACACTTGGTATC TTAATCTATCWGRAACTTWA ATCACMATTACCGGAAWA
 2961 TCTCATATGGTACTCTANAA AAAACCCANAATTATGTCAT TGGCTGTAATTTGTTAA KTTGCCCAARACCTTCTCAA
 3041 ASCTTAAATTTGACCAT TTTGACATGAGGRTACTCC KAATTTTAAACATGCCCT TARGGATTGIGARTGIGAAA
 3121 ARTCTTCAAACCTCTTCTYMG TAAACATGGCATAGCAATAA CCGAGCTTAAAGAACATGTCAM AAYCTCCATGGGAAARTTIG
 3201 TATGGGGGCTGGGAAAAA TGAAAATGCMGKGATGC ACGTTAACGAACTTGTCTC AAAAAGGTTWAATGARTTA
 3281 NAAACCTGGRWTGGGGTGA TAAATTAAATGTTTCCGAA ATGGGAAACACTTGAAGAAAAGA AGTCTCTAATGAAAGTGTG
 3361 CTCATATGGTACTCTANAA AAAACCCANAATTATGTCAT TGGCTGTAATTTGTTAA KTTGCCCAARACCTTCTCAA
 3441 GGGTTCTGAAACTAGAGAT TGTGTCATGTTGATGAAA AGANTGTTTACGTTGTTTC ATCAATCACCAGTGGGAAA
 3521 TAGATGATATTTCAGGGCY TACTGATGAGATGTCAGAG GTATGATAGGGTCTCTGG GCGGTAGAAGAAATAAGCAT
 3601 CCATTCTGTAATGAAATAA GATATYTGTGGAAATCAGAA CGAGAGCAAGTAACGGTCT TATGAAATTAAAGAAGTGG
 3681 ATTAGGTTGAATGAAATAT TTGGTGTGTTAGGGAGAA AAAGGAGGATAATCATATA TTAAATAGTGGGAGCAGCTTA
 3761 ACATCTTTAGGAGGTGAA TGTATGGAGGATGTAACAGCT TGGAGCATGGCTTCAGGTTGCTCA GATAGCATGGAGAATTGTA
 3841 TATGCACTGTTGATTCAA TTAACATCCCTCTCTCCCA ACAGGAGGAGGACAGAAGAT CAACTCACTTACCATCACTG
 3921 ATTGCGAGAAGCTTCGGAA GAGGAGTTGGGAGGAGCAGAGA GAGGACAAGAGTGTCTTATAA ACTCAAAATGCAAGATGCT
 4001 GAATCTGAGATATACGTAATGGCCAATCTGAAATTCA TCAAGTGAATTGAGTTGCTTC ATTCACTGAAACAGATTATA
 4081 TATATCAAACCTGCGAGTR TGGAGTCATTCTGACCCAT GAGTGGCCAATCTCACCCTC TTAAACAGATCGAGGAGG
 4161 GACAGCGATTTCGTCAGGAA CGGTTACGATTGCTACGGG 6TCGTTT

SEQ ID NO: 2

RLG1B
[Strand]

1	AACCGTTCGT	ACGAGAAATCG	CTGTCCCTCTC	CPTCCCTGTAA	TATAATGATA	AGAAAAAAATA	TGATTAAAGG
71	TITAAATCCA	AAATCCATTA	TTCCACCGGT	GATATGATGC	ACTAGCTGT	GTATGCACAA	ACAGTATTAT
141	AAATGCTAAC	CAAAACAGCA	GCTAAGAAC	AATATAAATA	ATGGTTTGAA	TGTCCTTTC	TCCGTACAC
211	CATTTCTTCC	AAATCCCTAT	CATTCTACAA	TACAAGTGT	CCCATTAGG	GTTTCACTA	TAGCAATGCC
281	CTGAAATCTCT	TGGTCTCGG	TTCTTGGGG	TGTCTTTGA	AAAGCTTGCT	TCIGAAGCCT	TGAAGAGGGT
351	TGCTTGGCTCC	AAAGTAATTG	ACAAGGAGCT	CGAGAAATG	AATAGCTCAT	GAATCAATAT	AAAAGCTCTG
421	CTCAATGATG	CTTCTCAGAA	GGAAATAAGT	AAGGAAGCTG	TTAAAGAATG	TTGAATGCT	CTTCACATT
491	TGCCCCACGA	CATAGATGAT	CTACTTTGGG	ATTTGGCAAC	CAAAGCTATC	CATCGTAAAGT	TCTCTGAGGA
561	ATACGGGGC	ACCATCAACA	AGGTACCGAA	GTAAATTCCA	TCTTGTCTCT	CTAGTTGTC	AAGTACTAAG
631	ATGCCCAACA	AGATACATAA	TATTACCGC	AAGTACAAG	AACTATTAGA	AGAGAGAAAT	ATCTTGGAT
701	TATGTGAAAT	TGGTAAAGC	CGAAAACCTC	GAATAGAAA	ATCAGAGACC	TCTINGCTAG	ATCCATCTAG
771	TATTGTTGGA	CCGACAGATG	ATAAGGAAGC	TTGCTTCTC	AAGCTATATG	AACCATGTGA	TAGAAACTTT
841	AGCAATCTTC	CNATGTTGG	TATGGGTGGG	TTAGATAAGA	CCACTTTAGG	TAGACTTTG	TATGATNAAA
911	TGCAAGTGAA	GGATCACTTC	GAACCTCAAGG	CGTGGGTTG	TGTTCTGAT	GAGTTTGATA	TCTTCGGTAT
981	AAGCAAAACC	ATTTTCAAT	CGATAGAGGG	GGGAACCAA	GAGTTTAAGG	ATTPAAATCT	GCTTCAGGTG
1051	GCTTTAAAGG	AGAAAATCTC	AAAGAAACGA	TTTCTTGTG	TCTCTGTGA	TGTATGGAGC	GAGACCTATA
1121	CTGATTGGGA	AATTCTAGAA	CGTCCATTTC	TAGCAGGAGC	ACCAGGAAGT	AAAGTAATCA	TCACAAACCG
1191	CAAGTGTGCG	TIGCTAAACC	AATTGGGTCA	TGATCAACCA	TACCAATTGT	CTGATTGTG	ACATGACAAT
1261	GCTCTATCTC	TATTCTGTCA	ACACGCATT	GGTGTAATAA	GCTTGTATT	ACATCCGATA	CTTAAACAC
1331	ATGGTGAAGG	TATTGTMGAA	AAATGTGATG	TTTGTCCATT	GGCTTGTATT	GCACCTGGGA	GGTTATTGAG
1401	GACAAAAAGA	GATGAGGAAG	AATGGAAGGA	ACTATTGAAT	AGTGAGATAT	GGAGGTTAGG	AAAGAGAGAT
1471	GAGATTATTC	CGGYTCTTAP	ACTAAGCTAT	AATGATCTTT	CTGCTCTTT	GAAGCAGTTG	TTTGCATATT
1541	GCTCCCTGT	CCCCAAAGAT	TATGTGTCA	ACAAGGAGAA	TTGATTTTA	TTATGGATGG	CAGAAGGGTT
1611	TTTGCACAAAT	AAAATACAA	ACAAGTCATA	GGAACCTTA	GNTCTTGAAT	ATTTTGACGA	CTTGTGTCA
1681	AGGTCACTTT	TTCAACATGC	ACTCGATGAC	AAATCGTTG	TTGTTGGCCA	CGACCTCATG	AATGACTTGG
1751	CCACATCTGT	TGCTGGAGAT	TATTTTTAA	GATTAGACAT	TGAATGAAA	AGGAAGCTT	TGGAAAAATA
1821	CCGACATATG	TCATTTGTTT	GTGAGAGTTA	CATGGTTTAC	AAAAGGTTG	AACCATTATA	AGGAGCTAAA
1891	AAATTGAGAA	CTTTCTTAGC	AATGCCCTGTT	GGGATGATAA	AAAGTTGGAC	AAACATTTTAC	TTATCAAATA
1961	AGGTCCCTGA	TGACTTACTT	CACGAATTAC	CATTTGAG	AGTTCTAAGT	TTGAGTTATC	TTAGCATCAA
2031	GGAGGTACCT	GAAATAATAG	GCAATTGAA	ACACTTGGG	TATCTTAATT	TATCACACAC	GAGTATCACA
2101	CATTTACCAAG	AAAATGTCTG	CAATCTTAC	AACTTACAAA	CATIGATCCT	TTGTTGGCTGT	TGTTTTATAA
2171	CCAAGTTTCC	CAACAATTC	TTAAAGCTTA	GAAATTACG	GCATTGGAC	ATTAGCGATA	CTCCCGGTTT
2241	GAAGAAGATG	TCTCTGGGGG	TTGGTGAATT	GAAGAACCTA	CACACYCTCT	CCAAGCTCAT	TATGGAGGT
2311	AAAAATGAC	TAACCGAGCT	TAAGAACTTA	CAAAATCTCC	ATG		

RLG1b - Diana
[Strand]

1 TACTACTACT AGAATTCCGGT GTTGGTAAGA CGACTCTAGC TAGACCTTTTG TATGAGGAAA TGCAAGGGAA
71 GGATCACTTC GAACTTAAGG CGTGGGTATG TGTTTCTGAT GAGTTTGATA TCTTCAATAT AAGCAAATT
141 ATCTTACAAT CGATAGGTGG TGGAAACCAA GAATTACGG ACTTAAACCT GCTTCGAGTA GCTTTAAAAG
211 AGAAGATCTC AAAGAAAAAGA TTCTCTCTTG TTCTTGATGA TGTTGGAGT GAAAGCTATA CCGATTGGGA
281 ATTINNTAGAA CGCCCCATTTC TTGCAGGGGC ACCTIGGAAGT AAGATTATTA TCACCCACCCG GAAGCTCTCA
351 TTGTTAAACA AACTCGGTGA CAATCAACCT TACAACCTTT CGGTTTGTC ACATGAGAAT GCTTTGTCTT
421 TATTCTGTCA GCATGCCATTG GGTGAAGATA ACTTCAATTIC ACATCCAACA CTAAACCAC ATGGCGDAGG
491 TATTGTGAA AAAATGTGATG GattGCCATT GGCAATTGTGCG ACATGATGAT GATG

SEQ ID 137

SEQ ID NO: 3
RIGIC
[Strand]

1 TCCCCTGCCAA CGTNTATCAT TCAGAAGNGC CCAAAGACCA NAGATNTGTT TAANGNTGNT TNTCAGAAGG
71 AAGTAATTGA TGAAGCTGTN AAAAGATGGC TGATTGATNT CCAACAATTG GCTTACGACA CTGANGACNA
141 ACTTGATGAT NTCCGAACAG AAGCTTATCA TCCTGAGTTG ATCCGTGAAA CTGGAGCTTC CNCCAGCAATG
211 GTAAGRAAGC TAATCCCCAG TTGTTGCAAC AGTTTCTCAC AAAGTAATAG GATGCATGCC AGGTTAGATG
281 ATATTGGCCG TAAGTRACAA GAACCTGGTAG AGGCAGAAAAA TAATCTTGGT TTAAGTGTGA TAACATAACGA
351 AAAACCCAAA ATTGAAAGAG ATGAGGGGTN TTGTTGAGAT GCAAGTGGTA TCATGGACG TGAAGATGAT
421 AAGAAAAAAAT TGCCTTCAGAA GCTGTTGGGG GATACTTATG AATCAAGTAG TCAAAACTTC AACATCGTGC
491 CCATAGTTGG TATGGGTGGG GTAGGTAAAA CAACTCTAGC TAGACTTTTG TATGATGAAA AAAAAGTGAA
561 GGATCACTTC GAACCTCAGGG TTGGGTTTG TGTTTCTGAT GAGTCAGTG TTCCCAATAT AAGCAGAGTT
631 ATCTATCAAT CTGTGACTCG TGAAAACAAA GAATTGCGAG ATTTAAATCT GCTICAAGAA GCCCTTAAAG
701 AGAAAACCTCA GAACAAAACTA TTCTTAATAG TTTTAGATGA TGTATGGTCT GAAAGCTATG GTGATTGGGA
771 GAAATTAGTG GGCCCATTTC ATGCTGGGAC TTCTGGAAGT AGAATAATCA TGACTACTCG GAAGGGACCAA
841 TTACTCAAAC AGCTGGGTTT TTCTCATGAA GACCCCTCTGC ATAGTATAGA CTCCCTGCCAA CGTCTATCAC
911 AAGAAGATGC TTGTCCTTG TTTCCTCAAC ACGCAATTGG TGTACCTAAC TTGATTTCAC ATCCAACACT
981 AAGGCCATAT GGGGAACAGT TTGTGAAAAA ATGTGGGGGA TTGCCCTTGG CCTTGT

SEQ ID NO:4

RLGID

[Strand]

1 CTTACCCCTTC TACCGAGATCC CTGTCCTCC TCGATCTCT TAACCATGCT TCCCAGAAGG AAGTNACTAA
71 TGAAGCCGTT AAAAGATGGC TGAATGATCT CCAACATTG GCTTATGACA TANACGACCT ACTTGATGAT
141 CTTGCCACAS AAAGCTATTTC NTICSTGAGTT GACCGANGAA GGTGGAGCCT CCACCCAGTAT GGTAAAGAAA
211 CTAATCCCAA GTTGTGAC AAGTTTCTCA CAAAGTTATA CGATGCATGC CAAGTTAGAT GATATTGCCA
281 CCAGGTTACA AGAACTGGTA GAGGCAAAAA ATAATCTTGG TTAAAGTGTG ATAACATATG AAAAGCCCAA
351 AATTGAAAGG TATGAGGCAT CTITGGTAGA CGAAAGTGGT ATTITGGAC GTTNAAGATGA TNAGAAAAAA
421 TTGATGGAGA AGCTGTGGA GGATAAAGAT GAATCCGGAG TCNAAACTTC AGCATCCTGC CCATAATTGG
491 TATGGGTGGA GTTGGCNAAA CAACTCTAGC TAGACTCTTC TTGATGAAA AGACAGTGAA GGATCACTTC
561 GAACTCAGGG CTGGGTTTG TGTTTCTGAT GAATTCAAGTA TTCTCAACAT AAGCAAAGTT ATCTTATCAAT
631 CTGTGACCGG GGAAAAGAAA GAGTTTGAAG ACTTTAAATCT GCTTCAGAA CCTCTTAGAG GGAAACTACA
701 AACACAAACTA TTCTAAATAG TTGATGGATGA TGATGGTCG GAAAGCTATG GTGATGGGA GAAATTAGTG
771 GGCCCCTTTC ATGCTGGGAC TTCTGGAAGT AGAATAATCA TGACTACTCG GAAGGAGCAA TTACTCAAAC
841 AGTTGGGTTT TTCTCATCAA GACCCCTGTC GTTGTATAGA CTCCCTGCAA CGTCTATCAC AAGATGATGC
911 TTGTCCTTGT TTGCTCAAC ACGCATTTGG TGWCCA

RLG1E
[Strand]

1 TCTAGCTAGA CTTTGTATG ACGAGATCCA AGAGAAGGAT CACTCGAAC TCAAGCCGTG GTTTGTGT
71 TCTGATGAGT TTGATATATT CAATATAAGC AAAATTATTT TCCAATCGAT AGGAGGTGGA AACCAAGAAT
141 TTAAGGACTT AAATCTCCTT CAAGTAGCTG TAAAAGAGAA GATTTCAAAG AAACGATTTC TACTTGTTC
211 TGATGATGTG TGGAGTGAAA GCTATGCCGA TTGGAAATT CTGGAACGCC CATTCTTGC AGGGGCAGCC
281 GGAAGTAAAA TTATCATGAC GACCCGGAAG CAGTCATTGC TAACCAAACG CGTTACAAG CAACTTACA
351 ACCTTCCGT TTTGTCACAT GACAGTGCTC TCTCTTATT CTGTCAGCAT CCATGGGTG AAGATAACTT
421 CGATTCACAT CCAACACTTA AACCACATGG CGAAGGCATT GTGAAAAAT GTGCT

SEQ ID NO:5

RLGLF
[Strand]

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1   ATTTTCNGCT CNAAACAAAN AAAAGCAATG GCTGAAATCT TTCTTTCNGC ATTCTAGACC AGTATTCTTT
71  GAAAAGNTGG CTTCTGAAGC CTTGAAGAAG ATCCCTCGCT TCCATCGGAT TGATTCTGAG CTCAAGAAC
141 TGAAAGGGTC ATTAATCCAG ATCAGATCTG TGCTTAATGA TGCTCTGAG AAGGAAATAA GTGATGAAQC
211 TGTTAAAGAA TGGCTGAATG GTCTCCAACA TTGCTCTTAC GACATAGACG ACCTACTTGA TGATTTGGCA
281 ACCGAAACTA TGCACTGTGA GTTGACCCAC GGATCTGGAG CCTCCACCAAG CTGTAAGAA AGATAATCCC
351 AACTTGTGTC ACAGATTTCT CACTAAGTAG TAAGATCGGT AACAGTTAG ATAATATTAC CATCAAGTTA
421 CAAGAACTGG TAGAGGAAAA AGATAATCTT GGCTTAAGTG TGAAAGGTGA AAGCCCCAAA CATAACAAAC
491 GAAGATTACA GACCTCTTTC GTAGATGCAT CTAGCATTT TGGTGTGAA GTGATAAGG ATGCATTGCT
561 CCATAAGCTG CTGGAGGATG AACCAAGTGA TAGAAACTTT AGCATCGTGC CAATAGTTGG TATGGGTGGT
631 GTGGGTAAGA CGACTCTAGC TAGACTTTTG TATGACGAGA TCCAAGAGAA GGATCACTTC GAACTCAAGG
701 CGTGGGTITG TGTTTCTGAT GAGTTTGATA TCTTCATAT AAGCAAAGTT ATCTTCCAAT CGATAGGTGG
771 TGGARACCA AATTTAAGG ACTTAARTCT CCTTCAAGTA CCTGTAAAAG AGAAGATTTC AAAGAAAACGA
841 TTCTCTYTTG TTCTGGATGA TTGTTGGAGT GAAAGCTATA CAGAATGGGA AATTCTAGCA CGTCCATTTC
911 TTGCAGGGGC ACCAGGAAGT AAGATTATCA TGACGACCCCG GAAGTTGTG TGCTAACCA AACTCGGTTA
981 CAATCAACCT TACAACCTTT CSGTTTGTG ACATGATAAT GCTYGTCTT TATCTGTCA SCAYGCATTG
1051 GGTGAAGATA ACTTCGATTC ACATCCAACA CTTAAACAC ASGGTGAAG TATGTGAA AAATGTGACG
1121 GTTTCACCATG GGCTTTRATT GCACTTGGGA GRTTGTGAR GACAAAAACA GATGAGGAAG AATGGARGA
1191 AGTGTGAAAT AGTGAATAT GGGGGTCAGG AAAGGGAGAT GAGATTGTTG CGGCTCTTAA ACTAAGCTAC
1261 AATGATCTCT CTGCCCTCTT GAAGAAGTTG TTGTCATACT GCTCTTGTG CCCAAAAGAC TATGTGTTGG
1331 ATTAAGGAGGA GTTGTGTTTG TTGTGGATGG CAGAAGGGTT TTTGCACCAA TCAACCCACAA GCAAGTCRAT
1401 GGAACGCTTG GGHCATGAAAG GTTTTGATGA ATIGTGTCA AGATCATTTT TICAAACATGC CCCTGATGCC
1471 AAATCGATGT TTGTGATGCA TGACCTGTATG AATGACTTGG CHACATCTGT TGCTGGAGAT TTTTTTCAAA
1541 GGATGGACAT TGAGATGAAAG AARGAATTAA GGAAGGAAGC TTGSAAGAG YAYCGCCATA TGTCAWTGT
1611 TTGTGAKGAT TACATGGTKK ACAAAAGTT CRAGCCATTS ACAAGGAGCT AG

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SEQ ID NO: 6

RLG1G
[Strand]

1 GTGAAGGATC ACTTCGAACT CAGGGCTTGG GTTTGTGTTT CTGATGAATT TAATATCCTC AATATAAGCA
71 AAGTAATTTA TCAATCTGTA ACCGGGGAAA AAAAGGAGTT TGAAGACTTA AATCTGCTTC AAGAAGCTCT
141 TAAAGAAAAA CTTTGGAAATC AGTTATTTCT AATAGTCTG GATGATGTGT GGTCIGAAAG CTATCGTGAT
211 TGGGAGAAAT TAGTGGGCC ATTTCCTCG GGGTCTCTG GAAGTATGAT TATCATGACA ACTCGGAAGG
281 AGCAAATGCC AAGAAAGCTG GGTTTCTTC ATCAAGACCC TTGCAAGGT CTATCACATG ACGATGCTTT
351 GTCTTGTGTT GCTAACACCG CATTGGTGT ACCA

SEQ ID NO:7

RLG1H
[Strand]

1 TCTACCTAGA CTTTGTATG AGGAATGCA AGGGAAAGGAT CACTTCGAAC TCAAGCCGTG GGTATGTGTT
71 TCTGATGAGT TTGATATCCTT CAATATAAGC AAAATTATCTT TACAATCGAT AGGTGGTGGAA ACCAAGAAAT
141 TTACGGACTT AAACCTGCTT CAAGTAGCTT TAAAAGAGAA GATCTCAAAG AAAAGATTTTC TTCTTGTTCT
211 TGATGATGTT TGGAGTGAAA GCTATACCGA TTGGGAAATT CTAGAACGCC CATTTCCTGC AGGGGCACCT
281 GGAAGTAAGA TTATTATCAC CACCCGGAAAG CTGTCATGT TAAACAAACT CGGTACAAAT CAACCTTACA
351 ACCTTCGGT TTGTCACAT GAGAAATGCTT TGTCCTTATT CTGTCAGCAT GCATTGGGTG AAGATAACTT
421 CAATTACACAT CCAACACTTA AACCAACATGG CGAAGGTATT GTTGAATAAT GTGAT

SEQ ID NO:8

RLGI
[Strand]

1 TCTAGCTAGA CTTGTGTATG ATGAGATGCA AGAGAAGGAT CACTTGAAC TCAAGGCCTG GGTATGTGTT
71 TCTGATGAGT TTGATATATT CAATATAAGC AAAATTATTTC TCCAATCGAT AGGAGGTGGG AACCAGAAAT
141 TTAACGGACTT AAACCTCCCTT CAAGTAGCTG TAAAAGAGAA GATTTAAAG AACGATTTC TTCTTGTTC
211 TGACCGACGTG TGGAGTGAAA GCTATGCCGA TTGGGAAATT TTGGAACGCC CATTCTTGC AGGGGCAGCC
281 GGAAGTAAAAA TTATCATGAC ACCCGAAAG CAGTCATTGC TAACCAAACG CGGTTACAAG CAACCTTACA
351 ACCTTTCCGT TTTGTACAT GACAGTGCTC TGTCCTTATT CTGTCAGCAT GCATTGGGTG AAGGTAACCT
421 CGATTCAACAT CCAACACTTA AACCACATGG CGAAGGCATT GTTGAAAAAT GTGCTGGATT GCCATTGGCA
491 TTGTCGACA

SEQ ID NO. 9

RLGLJ
[Strand]

1 TACTACTACT AGAATTCCGGT GTTGGTAAGA CGACCTCTAGC TAGACTTTTG TATGAGGAAA TCGCAAGGGAA
71 GGATCACTTC GAACTTAAGG CGTGGGTATG TGTTTCTGAT GAGTTTGATA TCTTCATAT AAGCRAAATT
141 ATCTTACAAT CGATAGGTGG TGAAACCCA GAATTTACGG ACTTAAACCT GCTTCGAGTA GCTTTAAAAG
211 AGAAGATCTC AAAGARAAGA TTTCCTCTTG TTCTTGATGA TGTTGGAGT GAAAGCTATA CCGATTGGGA
281 AATTNTAGAA CGCCCATTTTC TTGCAGGGC ACCTGGAAAGT AAGATTATTA TCACCCACCCG GAAGCTGTCA
351 TTGTTAAACA AACTCCGTTA CAATCAACCT TACAACCTTT CGGTTTGTTC ACATGAGAAT GCTTGTCTT
421 TATTCTGTCA CCATGCCATTG GGTGAAGATA ACTTCAATTAC ACATCCAACA CTAAACCAC ATGGCGAGG
491 TATTGTTGAA AAATGTGATG GATTGCCATT GGCATTGTGAC ATATGATGAT GATG

SEQ ID NO:10

RLGIA a.a.

IVTVRTR?LSLLHLLSYVIFS?I?PH?ILWLF.INFYSTCHFMSFSILLSFT.YLNVITINAYLFFFK.THIYR
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SILLRFHPNQYFQRMTDSYGVSEFAFRHCSLKEIINQMELLOCSLLMKGELYVK?MSA?LHPEPTTLSV
YHQTQNNGGSR?T?KS.RIDYFCPHGLTEERV.FIFL.?KNYRSIEFLHRQRSFTSQCFVKQFLIFLSFR.NS
SIATCNLQLLGPQICGGR.FNPHIHCKQ.FKSISVHPIHQHLLIEIIHASSISSTSILYSLLSY.TMAEIVLS
AFLTUVFEKLA?EALKKIVRSKRIESELKKLKETLDQIQDLLNDASQEVTKNEAVKRWLNDLQHLAYDID
DLLDD?ATEAV?RELTEEGGASSSMVRKLIPSCCTSFSQSNSRMHAKLDDIATRLQELVEAKNNLGLSVI
TYEKPKIERYEAISLVDSTVGRREDDKKKLLKLGDKDESGSQNFISIVPGMGGVGKTTLARLLYDEK
KVKDHFELRAWCVSDEFSPNISRVYQSVTGEKKEFEDLNLQEAKEKLRNQLFLIVLDDWSESY
GDWEKLVGPFLAGSPGSRIIMTRKEOQLRKLGFSHQDPLEGLSQDDALSLFAQHAFGVNPFDSHPTLR
PHGELFVKKCDGLPLALRTLGRLLRTKTDEEQWKELLDSEIWRLGKSDEIVPALRLSYNDSA?LKLLFA
YCSLFPKDYEFDKEELLWMAEGFLHQPT?NKSQRLGLEYF?ELLSRSFFQHAPN?KSLFVMHDLMD
LATFVAGEFFSRLDIEMKKEFRM?SLEKHRHMSFVCE?YIGYK?FEPFRGAKNLRTFLALSVGVVEDWK
MFYLSNKVLND?LQDPLLRLV?LI?L?I??VP??VGSM?HLRYLNLS?T?IHLPE??CNLYNLQTLIV
SGC?YLV?LPKTF?LKNL?HFDMR?TP?LKNMPL?IGELK?LQTLF?NIGIAITELKNL?NLHGK?CIGG
LGKMENAVGCTLSELVSKV?.?NW??G..ICFPKWEHLKKKSSMK.CLIMVL?KKP?IMSIGGIEFPN
WVGSLRVSETRDVFMVYEK?CFT.FHQSPSGK.MIFSG?TDEMWRGMIG?LGAVEEISIHCSCNEIRYLWE
SEAASKVLMNLKLDLGECENLVSLGEKKEDNNHNINGSSSLTSFRRLNVWRCNSLEHCRCPDSMENLY
MHMCDS?TSVSFPTGGGQKIKSLTITDCKLSEEEGLGRERTRVLINSKMQMLESVDIRNWPNLKSISEL
SCFIHLNRLYISNCPS?ESFPDHELPNLTS LTDERRGQRFSYERLRFDWPSF

SEQ ID NO:11

RLGIB a.a.

NRSYENRCPLLPVI...EKILKV.IQNPLFHR.YDALAWCKNSIINANQNSS.ETI.IMV.IVLSPYTHFFQIPII
HTYKCSHIRFLSLAMAEILGSAFFAVFFEKLASEALKRVACSKVIDKELEKLNSS.INIKALLNDASQKEIS
KEAVKEWLNALQHLPYDIDDLGLATKAIHRKFSEEGATINKVRKLIPSCFSSLSTKMRNKHNTS
KLQELLEERNNLGLCEIGESRKLRNRKSETS?LDPSSIVGRTDDKEALLKLYEPCDRNFSILPIVGMGGL
DKTTLGRLLYD?MQVKDHFELKAWCVSDEFDFGISKTIFESIEGGNQEKFDLNLLQVALKEKISKKRFL
VVLDVVWSESYTDWEILERPFLAGAPGSKVITTRKLSLLNQLGHHDQPYQLSDLSHDNALSFCQHAFG
VNSFDSDHPILKPHGEGIVEKCDGLPLALIALGRLLRTKRDEEEWELLNSEIWRLGKRDE!P?LRLSYND
LSASLKQLFAYCSLFPKDVFNKEKLILLWMAEGFLHNENTNKSMERL?LEYFDDILLSRSFFQHALDDKS
LFVWHDLMNDLATSVAGDYFLRLDIEMKKEALEKYRHMSFVCESYMVKRFEPFKGAKKLRTFLAMPV
GMIKSWTTFYLSNKVLDLLHELPLLRVLSLSYLSIKEVPEIIGNLKHLRYLNLSHTSITHLPENVCNLYN
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H

SEQ ID NO:12

RLG 1c a.a.

SRAT?IIQK?PKT?D?F????QKEVIDEAVKRWLID?QQLAYDT?D?LDD?ATEAIHRELIRETGAS?SMVRKLIPSCCTSFSQSNSRMRHARLDDIAAK?QELVEAKNNLGLSVITYEKPKIERDEA?LVDASGIIGRED
DKKKLLQKLLGDTYESSSQNFNIVPIVGMSGVGKTLARLLYDEKKVKDHFELRVWVVCVSDEFSVPNISRVIYQSVTGENKEFADLNLLQEALKEKLQNKLFLIVLDDWSESYGDWEKLVGPFHAGTSGSRIIMTTR
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SEQ ID NO:13

RLG ID

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LMEKLLEDKDESGVKLQHLPPIGMGGVG?TTLARLLFDEKTVKDHFELRAWCVSDEFSILNISKVIYQS
VTGEKKEFEDLNLLQEARLGKLQNKLFLIVLDDWSESYGDWEKLVGFHAGTSGSRIIMTRKEQLLK
QLGFSHQDPLRCIDSLSQRLSQDDALSLFAQHAFG?

SEQ ID NO:14

RLGIE

LARLLYDEMQEKDHFELKAWVCSVDEFDFNISKIIFQSIGGGNQEFKDLNLLQAVKEKISKKRFLVLD
DVWSESYADWEILERPFLAGAAGSKIIMTRKQSLLTKGYKQPYNLSVSHDSALSLFCQHALGEDNF
DSHPTLKPHGEGIVEKCA

SEQ ID No: 15

R L G I F

FSA?NK?KQWLKSFF?HSRPVFFEK?ASEALKKIARFHRIDSELKKLKRSLIQIRSVLNDASEKEISDEA
VKEWLNLQHLSYDIDDLDDLATETMHRELTTDLEPPPACCKDNPTCCTDFSLSSKMRNKLNDITIKL
QELVEEKDNLGLSVKGESPKNRRLQTSVDASSIIGREGDKDALLHKLLEDEPSDRNFSIVPIVGMGG
VGKTTLARLLYDEMQEKDHFELKAWCVSDEFDFNISKVIFQSIGGG?QEFKDLNLLQAVKEKISKKR
FL?VLDDWWSSESYTEWEILARPFLAGAPGSKIIMTRKLSLLTKLGYNQPYNLSVLSHDNALSLFCQHA
LGEDNFDSHPTLK?GESIVEKCDGLPLALIALGRILL?TKTDEEEWKEVLNSEIWGSGKGDEIVPALKLS
YNDLSASLKKLFAYCSLFPKDYVFDKEELLLWMAEGFLHQSTTSKSMERLGHEGFDELLSRSFQHAPD
AKSMFVMHDLMNDLATSVAGDFFSRMDIEMKKEFRKEAL?K?RHMS?VC?DYMV?KRF?P?TRS.

SEQ ID NO.:16

R LG1 G

VKDHFELRAWCVSDEFNILNISKVIYQSVTGEKKEFEDNLILQEALKEKLWNQLFLIVLDDVWSESYR
DWEKLVGPFFSGSPGSMIIMTRKEQLPRKLGFPHQDPLQGLSHDDALSLFAQHAFGVP

SEQ ID NO:17

RLG 1 +

LARLLYEEMQGKDHFELKAWCVSDEFDFIFNISKIILQSIGGGNQEFTDLNLLQVALKEKISKKRFLVLD
DVWSESYTDWEILERPFLAGAPGSKIITRKLSLLNKLGYNQPYNLSQLSHENALSLFCQHALGEDNFN
SHPTLKPHGEGLIVEKCD

SEQ ID NO: 18

R L G I F

LARLVYDEMQEKDHFELKAWCVSDEFDIFNISKIIFQSIGGGNQEFKDLNLLQAVKEKILKKRFLVLD
DWSESYADWEI?ERPFLAGAAGSKIMTRKQSLLTKGYKQPYNLSVSHDSALSLFCQHALGEGNF
DSHPTLKHGEGLIVEKCAGLPLALST

SEQ ID NO:19

RLG 15

EFGVGKTTLARILLYEEMQGKDHFELKAWCVSDEFDFNISKIILQSIGGGNQEFTDLNILLRVALKEKISK
KRFLVLDDWSESYTDWEI?ERPFLAGAPGSKIIITRKLSLLNKLGYNQPYNLSQLHENALSLFCQH
ALGEDNFNSHPTLKPHG?GIVEKCDGLPLALS

SEQ ID NO: 20

SEQ ID NO: 21
RLG 2A

1 TTINACCCAT AAATTCTCNA CCTGNNGGGA CAAAAACCTA AAAATGGCC ATAATGCNCA AATCAGNAAG
 71 GTTGANAAAAG CTCTAAGTTT TTINACCTCCA NCTGATGCNC NNCTCTCTA AAGTCANAT CCAAGCTTGC
 141 CCTCCAACTC TANCNCCCTC AAATGGCACCT CCTTCCTCTC AAAAGCACAC AAGAACACTT TCAAGCTCAA
 211 CCACACTCAC ACAAGCTCTA GAACNAGGGT TAGGGCACAT TTAGGGTTTT GCTCTCTGGA ATGGTGTCT
 281 AAAAGTGAGG CCATAATGTT CCTTATATAA GGCTCACTCC CACAATTAGG CTTTCAATCT GAACGTANTA
 351 CGCCCCAGTGT ACACATATGGT ACCTGCCAACG TACTCCGGTAG TCTCCGGTC AANAATACAC TCATGAGTAC
 421 GCGCAACCGTA CTTTCCCTTA CGCCCAGCGT ACTCAAAAGC CAAACATCTT TTCAAGGGAC TAATTTTGAC
 491 AACTTGAGGA AAGAAAAGGA TCAAAGANAT ATACTTGAT TCCGGGATGT TACATGAAG TTGANACCTT
 561 GGCTAAAAAA TAAATATGGT TGTTGGAAGCC GTTGGCTGAG CAAGCAACAA GGCTAAAATT CGTAATCTC
 631 AAATGGTGT ATTTCCTATT CCTTCTTATT ATTTTACTTG ATTTACGGGT AGTTTTTTT TCTTACAAAAA
 701 AATATTAAAG TTGATAAAAGT ATAGCCACTA AAATTGACTT TTTCACAAAC ATAATGTCAA ATGGTGGGTA
 771 TATGTATCAT GTTGTATTAN ATAATGAATA TGATGATNCT GTTCTATTAA ANCCAAAAAA ATTATCTAAT
 841 GATTTTTATAT TGGAAAACAA AGTTGTGATT TTINGCATPA TATAATCAA TCCNCTTTTG TNIGGGAGGT
 911 GGATAAATGT GGTAAATTTA NAACAAGTGT TTINACNTTG AAGGGTNTGG AAAGGGTGG AAAAGGTAAA
 981 ATGATAAAAT GTTACACAA ATGTGTGATC CGACTGAATA TNAATGTTAA GGATNATTGT ATAAATTTG
 1051 TGATATATAG TAAGCATAAA TATTTAGAAT TGTGACTAA ATTATATAATG TATNCNAACT GGATTGAAAC
 1121 ATTTCCTGATA TANATTGATA ATGAAAATGA GCAACCTAA CATACTTATC TTGGTAGTT TGTTTATTAT
 1191 ATTTCCTTATAA NAATATAGAA NCATCCCTT ATTTCCTAAC CATACTTGG ACAGCACTTGA ATAAATGGGA
 1261 AAAATGTACC TTGCTATTAA GCACAAAAAA ATTATAAAA TGTCATATTG TATTTAGCAC AAACAAAAAA
 1331 AAAAAACTTA TCCCTTTTGC ATTAGGTCAAC AAAGAAATAT AAAATGGAA ATGTTGTGCT ATTTCATGCA
 1401 CTAAGAGAAA CTATTTGCG TTATTAATG CGGTTAACCC AATAGAAAAA TGGAACTACA TTGTCATTAA
 1471 GCATGAAAAAA AAATAACTTT CCATTTTTG CATCCGGTCA CAATAATAGA AAAATGAAAG TACGTTGCTA
 1541 TTAGCGAAA CTAACTCTT TTTCCTTT TGGCATGTA TCATAATAA TAGACTAAA TACGTTAGTT
 1611 TTACATTTT AATACATTGA ATGTCTAAT CCACATGTT TCTATATAAA AGGGAAATGT ATTTTACTTA
 1681 TTCTTGATT TTGGCTCTC TTGGTAGTAC CCAAAACATC CCTCTATCCA TCTATCCAA CTAAAATAAT
 1751 GAAAACCTATA TTCTTCCCTAT TGTAGGGATG TTATAAATTG TGTAAATTGTT TTATGCAA AAAGTGTGTT
 1821 TTGTTAACTA GATTAACGAG ATTCAATTTC CAGCAATTTC GGAGAAGTTC ATCCATCTT TGGAATATGAA
 1891 GTGCAAGCCA AGTTCTTTAA CATGGAATAT GAGGTCCCTA TATGCTCAA AAATAGCAA TGAGAAATT
 1961 TTAAATTTGG ATCCCCATAA AAGAAAATTG TTAAATGGTT GTTTAATAT TGTCATATGT GTCCACCGGA
 2031 TGAGCCTAAT ATAGTTTAT AAGGGTAAA GGTGGGTTTG GTGGGGCCAT TTATCTTAT TATTTCTAA
 2101 AGTCAGAATT AAGTAAAAAA AATATAAGA TAAATACCTA AAGGATAAA ATCAATTAA TTGGACCAA
 2171 AGACCAAAGT TGTTAAGGGG CTGTTGTTT TTTTGTGAA GAGCTGTGCA ACCACTTTG TCTGCGCCG
 2241 ACAGACAAAGC TGCAAGACATA TGCCCTCGCA GAGTGTGTTG TTTTGAAG TGCGCAGACC AAAAAACG
 2311 CTGGCGGAGG TCTCTCTGGC GCATATATGT GTCACTGTCT TCAAAAGGTCT TCAGACCTCA TTTTAACCAA
 2381 AAAAAACGGT GACCAACGGT TTTTTTTTCT TTCTCTGTG GCTGAAAATG CATTTTTAAT
 2451 TTATGACA TGAAATTAAAG TTGAAAAAT TAAATTATTT CAACAGCTGT AGACGTTAAA AACAAACAGT
 2521 CTCTCTGTTG CAGACTGTGG ACATTTGGTC CACCTCTTCT ACCGCAGAGA CTTCAGATG TGGTCCGCG
 2591 ACTGCGACAA TTGGCTCTC AAATAAACAA ACATCACCTA ATTICACTAC ACCACACGGG CCTCCAAATGT
 2661 AACAAAAAAA AGGTGAAAC AAAGTTGCT ATTTCCTCAT ATCCAGGGG CATTTTATGTA AGAGTTATCT
 2731 AAATTTTGTG TCGGTAGATC AGTTCTCACAA TTAAACCGG GTAAAGTGTG TGTTGTACG CGGCACCTG
 2801 AAAGGTTGAA ANGTAACCTC CAAACCTGA CAANAACTGA TATGAAGTAT CAAGTTAGAG GTTCAATTGG
 2871 TGAAAGGATTC AGCTGGAGGT TGGGAAATCG AGCTTCCACT ATTAAGGTAA ATCCATAAC CCTAAATGT
 2941 GTTACCGCTCA TATATCAAAT TCGCTGTTT GTTGAATGAA AAAACCATGC TCAAAACCC AGTGTAAAGGC
 3011 AGGGTATATG ACATATTATG ATGTTACTGAT AACAATTTGATAAATTTG GTTGTACGTA AGTTAGGATT
 3081 CGTACTCTAA CCAATGTAA TAGTTTTGT GAGTCATATCT ATGTTATGG GGAATCACAT TAGCAACGGG
 3151 ATTGTACTAG TAATCGAAA AAGTCTTTA AATAATTTTT CTGTTATAA TTATGAAATA TTGTTAGCGA
 3221 CATCTAAAT TAAATAGAAAT GTATCTGATA TTGAAATTAAAT GTCCCTTAATG TGAACATAGA CCTTTCCAT
 3291 TTACTAATGC CTAAATTATA GTTCTCTAACT AATAAAATT AATTCCTGTT TTATGCTTCT AAGACAAATA
 3361 AAATCCATGA TTTCCTTTA AATAATCAA AAAATGACCA TAAATAAAAT AAAAATTAGG ATACCAAACC
 3431 CCCCGGGCAT GCCCAATGTC TAAATATCT GTGTCCTT GCTTCTCTCCTT TTGTCATATT
 3501 ATTCTGGAGA GTTTCATACA AGAAAATTTC AAGAAGAAAG CAAAGTCCA GTTATCTCT
 3571 TTCTCTAACTT ATGTTAAAC TTACAAGCAT TTTCACAGC ATCCATGGTT TTGTTGTAT GTTTTCAAA
 3641 TTGAAACTAG ATTGGGACTT TTGCCCCCTGTA TGATTCTATAA GATATTGCT GGAGTTGAGA TTGTTGAAAG
 3711 AAAGTGGTGA ATGAAAGAG CAAAGTGAATC CAGATATGTT ATGTTGATAA TATGATGATG AGATAGAGAT
 3781 ATGTTAAAC TGGCTAGAAA ATGTTTTAA TTGAAATTG AGGTGTTGAG ATGTTGAAAGA TACCAAGCTA
 3851 ATTAACCTAAAT AGTTATGCTA AATAGTATATA AAGAACACAA AACTCGTAGT TTGTTTTCTCA TGATTTCAAA
 3921 CCTCTCTGTA CCAAACTAAA TTATAACAA ATTGAATATC ATTCTCTGCA ATCAATTGTTA ACTTTTGT
 3991 TTATCTCAT GTCTAAATT CCCACAAGTT TATTTCTATA GTCATATTGG ATTATGAAAG GACTATT
 4061 ACCAATTCACA TTCTTACTTT ATGGCCAAAG CTAATACAAT CCGACTAAAC TAAAGGATTC TAGGATGCA

*SEQ 50 NO: 21
RLG 2A cont.*

4131 ATAGTTTGCT CCCCCATTAT AGATTTCTAT CTAAATTGTC TATTGTACTA ATTAGGTGC CACCACAAAGT
 4201 AAATTCCCTGA AATGGATGTC GTTAATGCCA TTCTTAAACC AGTTGTCGAG ACTCTCATGG TACCCGTTAA
 4271 GAAACACATA GGGTACCTCA TTCTCGCAG GCAATATATG AGGGAAATGG GTATCAAAT GAGGGGATIG
 4341 AAIGCTACAA GACTTGGTGT CGAAGAGCAC GTGAACCGGA ACATAAGCAA CCAGCTTGAG GTTCCAGCCC
 4411 AAIGTCAGGGG TTGGTTTGAA GAAGTAGGAA AGATCAATGC AAAAGTGGAA AATMCCCTA CCGATGTMGG
 4481 CAGTTGTTTC AATCTTAAGG TTAGACACCG GTCTGGAAAG AGAGCTTCCA AGATAATTGA CGACATCQAC
 4551 AGTGTCTATGA GAGAACACTC TATCATCATT TGAAATGTC ATTCCATTCC TTAGGAAGA ATTGATTCAC
 4621 CGAAAGCATC CACCTCAATA CCATCAACCG ATCATCATGA TGAGTTCCAG TCAAGAGAGC AAACTTTCAC
 4691 AGAACGACTA AACGCACTCG ATCCCAACCA CAAATCCCAC ATGATAGCCT TATGGGAAT GGGGGAGTG
 4761 GGGAAAGACGA CAATGATGCCA TCGGCTCAA AAGGTGTAAG AGAAAAGAA AATGTTTAAT TTTATAATTG
 4831 AGGGGGTTGT AGGGAAAAAA ACAGACCCCA TTGCTATCA ATCAGCTGTA CGAGATTAC TAGGTATAGA
 4901 GCTCAATGAA AAAACTAAAC CAGCAAGAAC TGAGAACGTT CGGAATGGT TTGTTGGACAA TTCTGGTGGT
 4971 AAGAAGATCC TAGTCATACT CGACGATGTA TGGCAGTTG TGGATCTGAA TGATATTGGT TTAAGTCCCT
 5041 TACCAATATCA AGGTGTCGAC TTCAAGGTGT TGTGACATC ACCAGACAAA GATGTTTGCA CTGAGATGGG
 5111 AGCTGAGGT AATTCACCT TTAAATGTAAG AATGTTATAA GAAACAGAAC CACAAAGTTT ATTCCACCAA
 5181 TTATATGAAA TTGGGATGTA TTGTTGATCTT GAGCTCCATA ATATAGGAGT GAATATTGTA AGGAAGTGTG
 5251 GGGGTCTTAC CATGCCATA AAAACCATGG CGTGTACTCT TAGAGGAAA ACCAAGGATG CATGGAAGAA
 5321 TGCACTTCTT CGTTTAGAGC ACTATGACAT TGAAAATATT GTTAATGGAG TTTTTAAAT GAGTTACGAC
 5391 AATCTCCAAAG ATGAGGAGAC TAAATCCACC TTTTCTTCTT GTGGAATGTA TCCCGAARAC TTGATATTIC
 5461 TTACCGAGGA GTTGGTGGAG TATGGATGGG GGTGAAATT ATTAAAAAA NTGTTACTA TAGGAGAAC
 5531 AAGAACCCAGG CTCAACACAT GCAATGGGGC GCTCATCATC ACAAAATTG TGATGGAGT TGATGATGTT
 5601 AGGTGCTATCA AGATGATGTA TCCTGTTGCT GCTTTGTTT TGATGATGTA TTCTAAAGTC GACCATGCTT
 5671 CCATTGTCAA CCATGTAAT ACACAGAGT GGCAATGCAGA TAATATGCAC GACTCTTGTAAAGACTTTC
 5741 ATTAACATGTC AAGGGTATGT CTAAGTTTC TACAGACCTG AAGTTCCAA ACCTCTCCAT TTGAAACTT
 5811 ATGCAATGAG ATATATCATGTT GAGTTTCCC AAAAACTTTT ATGAGAAAT GGAGAAGCTT GAGGTATAT
 5881 CCTATGATAA ATGAAAATAT CCATTGCTT CTCATCATCACC TCAATGTTCC GTCAACCTTC GGGTTTCA
 5951 TCTACATAAA TGCTCGTTAG TGATGTTGTA CTGCTCTGT ATTGGAAATC TGCGAATCT AGAAGTGT
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 6161 CTATAATGACA GTGGTTGATC GAGGTGAAAG CGCGATTAGC CTACAGATG ATAATCTGAA GGAGATGGCA
 6231 GAGCGITCAA AAGATATTTA TGCAATTAGAA CTTGAGTTCT TTGAAAACGA TGCTCAACCA AAGAATATGT
 6301 CATTGAGAA GCTACAACCA TTCCAGATCT CAGTGGGGCG CTATTTATAT GGAGATTCCA TAAAGAGTAG
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 6441 TTGTTTAAAGA AAACAGAGGT TTGTTGTTA AGTGTGGAG ATATGAATG TCTTGAAGAT ATTGAGGTTA
 6511 AGTCATCTTC AACAATCTT CAAATCTTCTT CGTCAACAA TTTAAGAGTC CTGTCGTTT CAAAGTGTGC
 6581 AGAGTTGAAA CACTCTTCA CACCTGGTGT TGCAAAACACT TTAAAAAAAGC TTGAGCATCT TGAAGTTAC
 6651 AAATGTGATA ATATGGAAGA ACTCATACATG AGCAGGGTA GTGAGAAGA GACGATTACA TTCCCCAACG
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 6791 ACCACAACTC ATGGAGTTG AACTTGACGA CATTCCAGGT TTCAACAGCA TATATCCCCAT GAAAAGTTT
 6861 GAAACATTGAA TTGTTGTTGAA GGAAGAGGTA AATATAAAATT TTAAATGCTA ATACATTACA AAGGATCTT
 6931 TCAGTTAAAT CTTTCAAAAT ATATGTAAT TTGATTGTT GGGTATTAT TTGTTGGATGG GACTTAAAT
 7001 AAATGTTTCTT CTGATTCTCA AGTTAGAGAA ACTGATGTT AGTAGTATGT GGAATCTGAA
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 7141 GATAAGCTTG TGAATTGTTT CTCGCACAAAG CCCATATCTC TGCTGCATCA TCTTGAAGAG CTAAAGTC
 7211 AGAATTGTTG TTCCATTGAA TCGTTATTC ACATCCATTG GGATTTGTT GTGCAACTG GAGATGAATA
 7281 CAACAAACAGT GTGTTAAGAA TTATTAAGT GATCAGTTGT GATAAGCTTG TGAATCTCTT TCCACACAA
 7351 CCCATGCTCA TACTGCATCA TCTTGAAGAG CTGAGTGCAGA AGAATGTTG TCCATTGAA TCGTTATICA
 7421 ACATTGACTT GGATTGTTG GGTGCAATTG GGCAAGAAGA CAACAGCATC AGCTTAAGAA ACATCAAAGT
 7491 GGAGAATTTC GAGAGGCTAA GANAGGTGAG GAGGATTTAAAGGTTGGAGATA ACTCTCGTCC CCTTGTTCAT
 7561 GGCTTTCAAT CTGTTGAAAG CATAAGGGT ACNAAAATGTN AGAAGTTTAG AAATGTATTIC ACACCTACCA
 7631 CCACAAAATT TAATCTGGGG GCACCTTTGG AGATTCATG AGATGACTGC GGAGAAAACA GGGGAAATGA
 7701 CGAACCGGAA GAGAGTACCC ATGAGCAAGA CGAGCTTAAAG ATTCAATTCTT CACTGTCCTTA ATTAAATGATT
 7771 AGCTTCTGCT TTGTTGATA AAAAGGGCA AACCATTTT ATGACTTAAT GTAGCAATAC AAGTCATGTA
 7841 TAAGAGTGTAC CAACTCTTIT TTATTTATAA AATGACTACA AAATTTTTT TTCTTATTAGA GATCAATGAT
 7911 AAATGTGACT AATTTTCTAT CACCTAATT TAGTTGATAA ATCTTATATAA ATGTCACTAG TTACTTTCA
 7981 GTAAAATAAC AAATTTAATC AATTATCAC AAAAGACATC AACTAAAAAATCCACAAAC CCGTAATAAT
 8051 TTAAAATAAA AGGATTAAAC ATCTAATACG AACATTTTT TTCTAAACA TGTTTGGAC CAAATATCAC
 8121 CAGCAACTCA AGTTGGAAAT CGATTCACT TAAACATTGA CCAGCATAAT TAGATAGATG AGAGTTGAAG
 8191 CTAAAGTGCCT TATATAAGTT CGTTTCATCT TTTTCTTGA TCTTGTAGC AAGTTGAATG ATTTCCTCT

RLG 2 A cont.

8261 TCAAAATTGA TAAAAATCTA CATTATAAAG AGACTAGCCT GAAAAAAAAT GGTCAGGTG GGTCCTGGGT
 8331 TCTCGTAGAT CAAGATGGAA GGGGAGAGTA TGATTTCAA GACACAACAC ATCCCTCATI TTATTTATTT
 8401 ATTATTATTA TTATTTTTTG ATATCTTGTG CATATTGTT ACAGATATGT GAGGTCTATT AATCTTTATA
 8471 AATATATAAA AAAATAATA ACATAAATG GAAAATTTAA TAAAGAATAA ATTAATAAGG GCACAATAGT
 8541 CTTTTAGGT AAGACAAGGA CCAAACACGC AACAAAAATA AACAGTAGGG ACCATCCGAT TTAAAAAAA
 8611 TAATTAGGGA CCAAAAACAT AAATTCCCCC AAACCATTAGG GACCATTACG GTAACTTACT CTTACTTTTC
 8681 GTTTTGTCA TATTGGGTA ACTATTTTGT TTGTACACAT CTAGGTAAACG AACTGTTGA AGTGTTCCCA
 8751 TTAGGATGT GACCTACTAC AACCGATCAT AATAGTCATA TGTGAACACT TCCAACAACT TTATTACTTA
 8821 GGTGTTACA AAAAACAT AGTTACCATG ATGTGAACAT ACTGAAAAAT TAATTACCTT AGCAAGTTAT
 8891 TTCCCTTTT AGGTGTATG GAAACAGTTC CGTGAGACCG TGACTGGAT GGTAGATAAA TTAGTAAAC
 8961 TTAACCCCTTC AATTAACCTA CCTTTTTCTT ATTAACCTAA TTCAACCTA AATTCCTGATT CTGTGTTGAA
 9031 AGTAAGTTGC ATCTTTATT TTGTATTATC TTGTTGCATA GGATCCTTAG CATCTTTAA TAATTTATTT
 9101 GAAGGTGAAA GATCCAACTA TTTTAAATCT GTGGCATTTC TCCATCATTT GCAACTGTTT CTGTAAAAAA
 9171 AAATACCTA AATCAAATA ACCATTTICA AATCCAAAT TATAAGAGAG AATTTAAAT GGACATGGAA
 9241 TCATAAATCA TTAACACAGT TCAGTAAACA AGTTGCTATT TACATTCTT GCTGTCAGA TTGAAATTCT
 9311 ATCAGAGAAA GAGACATTAC AAGAACCCAC TGACAGTATT TCTAATGTT TATTCCCATC CTGCTCATG
 9381 CACTCTTTTC ATAACCTCA GAAACTTATA TTGAACAGAG TTAAAGGAGT GGAGGTGGTG TTGAGATAG
 9451 AGAGTGAGAG TCCAACAACT AGAGAATTGG CAAACACTCA CCATAACCAA CAACACCTA TTATACCTCC
 9521 CAACCTTCAG GAATTGATTG TATGGAATAT GGACAAACATG AGTCATGTTG GGAAGTGCAG CAACTGGAAT
 9591 AAATTCTTCA CTCTTCCAAA ACAACAACTA GAATCCCCAT TCCACAACT CACAACCCATA AAAATTATGT
 9661 ATTCGAAAAG CATTAAAGTAC TTGTTTCG CTCATGCC AGAACCTCTT TCCAACCTAA AGCATATCAA
 9731 GATAAGGAGAG TGTGATGGTA TTGGAGAAGT TGTTCACAC AGAGATGATG AGGATGAAGA AATGACTACA
 9801 TTACACCTTA CCCACACAAAC CACCACTTGT TTCCCTAGTC TTGATTCTCT CACTCTAACT TTCTGGAGA
 9871 ATCTGAAGTG TATTGGTGGA GGTGGTCCCA AGGATGAAGG GAGCAATGAA ATATCTTCA ATAATACCAC
 9941 TCGAACTACT GCTGTTCTTG ATCAATTGGA GTTATGTTT GTACATATTIC AATTATTTAT TTATTTCTT
 10011 TTTTTATTG CAATATTCTA TAAATATAAC ATTATTAACC CACTTACTA AGATAATAAT TACCTAGAGG
 10081 GATGGATGCT ATGACACAGC TGCTACACTC CAGAAACTCT AGTAAGGCCA GTTATGGAAG TTCAATAAAAA
 10151 TGATAATGGC ATCTTTTGAT GGGTAATATA GGCAATTAA GTTTTATTTC TGTAAAGCA GTATTTAGCA
 10221 AGTACTGGCC AGTACGGAGAG CAGAATATCA CCTTTTGTGA AAATCTGGTC ATTTGACCCCA GAATTTAGTT
 10291 AAATGTAAACA TTTTAGATAT CAGGGGTCT CAGGTGACAG ATATGTAGA ATAGAACAAT ATATAATATC
 10361 ACCCTTAACT ATTCTTCTA AGGTTATTCT GTTAAATATG TGCCTTCTTG TTTCATNGA ATINGCATTIC
 10431 GTATATTATA GGTTAAAG TGATTTTNTC TCAATAAAT CCCGAAATTAA ATTAAAAAAA AAAAACAAA
 10501 AGTACATTTTG TGATGTCGAG ACCACTGGTA TCACITAGTA TATAAAAGC TTGATTITGA ATTAACCTTC
 10571 TTATACAAAAA GTTGTCTATA TAGTTTAAAT AGTTTACAT CATTCTTCA TGIGGTGTGAG CAGTGTCTG
 10641 AAGCAGGTGG TGTCTTGG AGCTTATGCC AATACGCTAG AGAGATGAGA ATAGAATTCT GCAATGCATT
 10711 GTCAAGTGTG ATTCCATGTT ATGCAGCAGG ACAATGCAA AAGCTGAAGG AGAGGACAGC GATTCTCGTA
 10781 CGAACGGTTA CGATTCGACT GCCCGTCGTT TTACA

SEQ ID NO: 21

R L G I A a.a.

MDVVNAILKPVVELMVPVKKHIGYLISCRQYMRMGKMRGLNATRLGVEEHVNRRNISNQLEVPAQV
RGWFEVGKINAKVENFPSDVGSCFNLKVRHGVGKRASKIIEDIDSVMREHSIIWNDHSIPLGRIDSTK
ASTSIPSTDHHDEFQSREQTTEALNALDPNHKSHMIALWMGGVGKTTMMHRLKKVKEKKMFNFII
EAVVGEKTDPIAQSAVADYLGIELNEKTTPARTEKLRKWPVDNSGGKKILVILDDWWQFVDLNDIGLS
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CGGLPIAIKTMACTLRGKSKDAWKNALLRLEHYDIENIVNGVFKMSYDNLQDEETKSTFLLCGMYPE?FD
ILTEELVRYGWGLKLFFK?YTIGEARTRLNTCIERLIHTNILLMEVDDVRCIKMHDLVRAFVLDMSKVEH
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VISYDKMKYPLLPPSPQCSVNLRVFHLLKCSLVMFDCSCIGNLSNLEVLFSFADSADRLPSTIGKLKKLR
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NMSFEKLORFQISVGRYLYGDSIKSRHSYENTLKVLEKGELLEARMNELFKTEVLCLSVGDMNDLEDIE
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GDEYNNSGVRIIKVISCDKLVNLFPHPNPMSSILHHLEEEVENCGSIESLFNIDLDCAGAIGQEDNSISLRNI
KVENLGKLR?VWRIKGGDNRPLVHGQSVESIRVTKC?KFRNVFTPTTNFNLGALLEISIDDCGENR
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RELVTTHNQQQPIILPNLQELILWNMDNMSSHWWKCSNWNKFTLPKQQSESPFHNLTTIKIMYCKSIKY
LFSPLMAELLSNLKHIIKIRECDGIGEVVSNRDDEDEEMTTFTSTHTTTLFPSLDSLTLFLENLKCIIGGG
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SEQ ID NO: 22

RLG 2B

SEQ ID NO: 23

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 71 ACAATGTAGA ATAATACTGG TATAATTAAAT TATATAAGT TATTAGCTG AAATCTTGAG GCTACTATAA
 141 TTAAATTATC ATAATTGAA AATCATAAA TTGTATTCGA TTGTATTTA TTGTATCAGA TAATTAAATAA
 211 TATGTGGGCC ACACAAATCC ACATCATCAG ACACCCACC TTATGTCGG CTACCTCACC ACTTGCATGA
 281 TCCCCACATC TTCCCAACCC CACCGACGAC TTGGGTCTC CTTAATATAT CAATTATTT CTGTAAAGTAT
 351 TTATTGTTGT AAATGTGTA TGTCATTATA CCTTTTTCT AATATAATTA GAACATAAA TTAAATATGA
 421 AATTCACACT CGTTTTCATTG TTGCAATTAA AAAAAGACT GTACTGTTG CAATATTATA TTATAACCT
 491 GATTAATTAA TTAAAGCTTA ATTGCATAAA TTGCAATTAGG TTGTATTTT TTGTATTTA GGGAGGGTGA
 561 GGGTCACCGG GAATCAAAGC ACTTATGTA AAGCAGGGGA ATACAAAAA ATTACTCGA AACAAATTIT
 631 ATTCAATTAA AGTGAGATAA TAATGTTCTG ATTAGATTAT GAGAAGTAGG AGATTTAAGT GATATACTCC
 701 ATTAAAGA AATTGCAATT TTAATTITGG ATCTCTGTG ATGACAAAAA TTAACTCGTG ACAGGTTATA
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 841 ATTAAAGG GGGTAAACAA TATCAAAATA CTTGATAAGT AATTATATAA ATATGCATTI ACCCTCTAA
 911 AGAAAATGCT ACTAAGCTTG GACCATCTCA GAATTACAAT CATACTCTC CCCTAAAAA AGATTCGTAT
 981 ATATCATGTC ATTGGCATT CATTCTTT TCACAACTCA TAGTCTATT CTCAAAAAAT TCGAGTTCTC
 1051 GTATTGTTAA GGAAAGATCAG AAGAGACTGT TCACACAGGT ACTCTCTTT ATTATATGAT TCACATCTAT
 1121 ATATGTTATT GTTCTCTTC TTATGGTT CGTCAGCTTA ATTCGCTTG CTGATTAAA TTCTCTCACT
 1191 TTCTTCACCG GATTTTTTA ATTATGGTT TTGTAAATGAA CAATGGTGA AGGAAAGAAA CATGGGAGTC
 1261 TTTCTCTAAG TAAACCTAGA TACTTAGTT ATAAGGGTAT ATGCTAAAT GAACATATGCC CATTCACTT
 1331 TGCCCTTTCT TTTACTTTT AGTTTTAGA ATCCAAGTT TCATPATGTG CTGGATGTGT GAGAAGAATA
 1401 GGCATTAAGA AGGTAAAGGA CGTACATAAA ATTGATGTT TAGTGAATGT TTCTTGATAT CATTATTTT
 1471 ACTCTCTAA AAACATATA GATCAACAC AAATGCTAC TTGTGTAGT AACAACCTCG ACTTAATAAT
 1541 GTTAATAATC AAGATTCCT TGATTCAC TATTTCCTAA CGAACACAAGC TCACTAAAAA CTCATATTC
 1611 TTGAGCTG AGTGGTTTAT ATTGGGGTT TTACATTAA TTCTTGTCG ATGAATGTGA AAATAGACTG
 1681 CTTATTGATT TTGCTGTTT CATTGAGTT ATTTCTATT TTACTACCTT ACAAATGCT CAGTGATAGA
 1751 TTCTCTAA TTGCTAACTT CCGTCTGCT TAAATATGTA GGAGCTACTA AAAGCAAAAA TTGAGCAA
 1821 TGTCGGACCC AACGGGATT GCTGGTGCCT TTATTAACCC AATGCTCAG ACGGCCTTGG TTCCCGTTAC
 1891 GGACCATGTA GGCTACATGA TTTCCTGCAG AAAATATGTC AGGGTCATGC AGATGAAAAT GACAGAGTTG
 1961 AATACCTCAA GAATCAGTGT AGAGGAACAC ATTAGCCGG ACACAAAGAA TCATCTTCAG TTCCATCTCA
 2031 AACTAAGGAA TGGTGGACCA AAGTGAAGG GTCAGAGCA AATGTTGAAA ACTTCCGAT TGATGTCATC
 2101 ACTTGTGTTA GTCTCAGGAT CAGGCACAGA CTGGACAGA AAGCNCTAA GATAACTGAG CAGATTGAAA
 2171 GTCTAACGAG ACAACTCTCC CTGATCAGTT GGACTGATGA TCCAGTTCTY CTAGGAAGAG TTGGTCCAT
 2241 GAATGCCATCC ACCTCTGCT CATTAACTGTA TGATTCTCCA TCAAGAGAGA AAACCTTTAC ACAAGCACTA
 2311 ATAGCTCTCG AACCCAAACCA AAAATCTCAC AGGTCTGTT GACATCACCG GACTCACAG GGGTGGAGTG
 2381 GAATGATGCA AAGGCTGAAG AAGGCTGTT AAGAAAAGA ATTGTATAAT TATATTGTTG GGGCAGTTAT
 2451 AKGGGAAAGG ACGGACCCCT TTGCAATTCA AGAACGCTATA CGACATTACC TCGGTATACA ACTCAATGAA
 2521 AAAACTAACG CAGCAAGAGC TGATAAGCTT CGTGAATGGT TCAAAAGAA TTCAAGATGGA GGTAAGACTA
 2591 AGTCTCTCAT AGTACTCTGAC GATGTTTGC AATTTGTTGA TCTTGAAGAT ATTGGGTTAA GTCCCTTTCC
 2661 AAATCCTAGGT GTGCACTTCA AGGTCTGTT GACATCACCG GACTCACAG GGGTGGAGTG
 2731 GAAGCTTATT CAATTATTA CCGGGGCTT CTAACCTGAAG CAGAACGCTCA AGTCTGTC CAAACATTG
 2801 TAGAACTTC TGAGCCGAG CTCCAGAAGA TAGGAGAGGA TATCGTAAGG AAGTGTGCG GTCTACCTAT
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 3081 GATGAGGTAT GGATGGGCT TGAAGCTATT TGATGAGTT TATGCAATT GAGAAGCAAG AACCAAGCTC
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 3221 TGCATGATCT GTTGTGTTGG GTATGTTTC TGAGTCAGG CAATCTCTATA TTGTCACCA
 3291 TTGTTATATT CCGGGGTTGC CTGATGAAAAA TGATGATGC GTGCACTCTT GCAAAAGAA TTCAATTAA
 3361 TGCAGGGTGA TGATTGAGAT CTCAGTACG CTCACATTTC CTAAACTAAC GATTTTGAAA TTATGCTGAT
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 3501 TAAAATGAAG TACCCATTGC TTCTTTGGC ACCTCGATGC TCCACCAACA TCGGGTGCT TCATCTCACT
 3571 GAATGTTCTA AATAGATGTT TGATGCTCT TCTATCGAA ATCTATCGAA TCTCGAAGTC CTGAGCTTIG
 3641 CAATCTCA CATTGAATGG TTACCTCTCA CAGTCAGAAA TTAAAGAAG CTAAAGTTAC TTGATCTGAG
 3711 ATTCTGATG GGTCTCTCTA TAGAACAGGG TGTCCTGAAA AGTCTTGTCA ACTTGTGAG ATTATATATT
 3781 GGAGATCTCAT CTGGTTTAT AGATGATAAC TGCAATGAGA TGCGAGAGCG TCTTACAAC CTTCTGCT
 3851 TAGAACTCCG GTTCTTTAA AACAAAGCTG AAGTAAAAAA TATGTCTATT GAGAATCTTG AACGATTCAA
 3921 GATCTCTG GGTGCTCTT TTGATGAAAAA TATCAATATG AGTAGCCACT CATACGAAAAA CATGTTGCAA
 3991 TTGGTGCACCA ACAAAAGGTGA TGATTAGAC TCTAAACTTA ATGGGTTATT TTGAAAACA GAGGTGCTT
 4061 TTGAAAGTGT GCATGGCAAG AATGATCTG AAGATGTTGA GGTGAAGTCG ACACATCTA CTCAGCTCTC

RLG 2B cont.
SEQ ID NO: 23

4131 TTTCATTCAG AATTTAAAAG TTCTTATTAT TTCAAAGTGT GTAGAGTTGA GATACCTTT CAAACTCAAT
 4201 CTTGCAAACA CTTTGTCAAG ACTTGACCAT CTAGAAGTT GTGAATGTGA GAATATGGAA GAACTCATAC
 4271 ATACTGGAAT TGGGGGTGTTG GGAGAAGAGA CAATTACTT CCCTAACGCTG AGTTTTTAT CTTTGAGTC
 4341 ACTACCGAAG TTATCAAGTT TGTCACCATTA TTGCAACATA ATTGGCTAC CACATCTCGT AGACTTGATA
 4411 CTTAAGGGCA TTCCAGGTTT CACAGTCATT TATCCGCAGA ACAAGTTGCG AACATCTAGT TTGTTGAAGG
 4481 AAGGGGTAGA TATATGTCT TTAGTGTAA ACAATTAAA TAATATTTTC ACCAAATTTC TCATAATATA
 4551 TCTGTAATTG GATGTTATG TGTTGTTATG TTATATGTCG GCTATTAAGG CATGATTATT TTGAGGTG
 4621 TGATTCTAA GTTGGAGACA CTTCAAGTTG ATGACATGGG GAACCTAGAA GAATATGGC TTGTTGAAC
 4691 TAGTGGAGGT GAGAAAGTTA AGTTGAGAGC GATTAAGTG AGTAGTGTG ATAGCTGTG GAATCTATT
 4761 CGCGCAATC CCATGTCCT GTTGCATCAT TTGGAAGAGC TTACAGTCGA GAATTGCGGT TCCATTGAGT
 4831 CGTTATTCAA CATTTGACTTG GATTGTTGCG GTGCAATTGG AGAAGAAGAC AACAAAGAGCC TCCTTAAGAAG
 4901 CATCAACGTG GAGAATTAGG GAAAGCTTAAG AGAGGTTGAGG AGGATAAAGG GTGCAAGATAA CTCTGATCTC
 4971 ATCAACGGTT TTCAAGCTG TGAAAGCTAA AAGATTGAAA ATGTAAGAG GTTTAGAAAT ATATTCACAC
 5041 CTATCACCCG CAATTTTAT CTGGAGGCC TTTTGGAGAT TCAGATAGAA GTTGGCGGAG GAAATCACGA
 5111 ATCAGAAGAG CAGGTAAACGC TTTCATTTT ACCTTCTAA TTATTAAGG ACTAAGCTCC TGTTTTTG
 5181 ATAATAAAGA GGTGGGATGA CTAAACTTGG GCATCACAAAT TGCAACAAAA TGTTACAAAC CATGAAACGT
 5251 TCAAAACATT TCTTGAATT AGGTTCAAT ACAAGTCATT TAAATATG GCCTAAATTTC TTGTTATATT
 5321 TATGTTATCAA CATGATTTTT CATTAGAGAT CATTATTATA ATAGTAAGTT TAAAGCAATT TAAATCAGAA
 5391 CTAATTCTAA CTTTAGCTAA TAAATCGTTA TAAATGTTA TAAATTACTT TTAGTGAATT AAGCAACGGA
 5461 TTAA-TAAGT TAACAACCTA AATGTCATT CCTAACAAAA AAAACTTTGG TTCAGAAAAAA CCGCAATTCA
 5531 AGATAACTAA AATAAAAATA TTGACATTC ACTAAAGAGCA TTGTTTTTTC TAAATATGAT TGCAATGAA
 5601 TAAACACTTAA ATTATACAG AAAATTCTTT TATATATGTT ATCAAATTG TAAACATTGAA AATTTGGATAT
 5671 GTTAA-TTAAAC CGTTTATAAT TCTGGTATCA CAAAGGGATA TATAATAAA TATTATTTTC TGTTAGTCATT
 5741 TGTAA-TTGTG TGTTTTATA ACCCGTGGGA ACCATGAGIT CTAAAATTAG TAAACTTTTC ATAATAAAAAA
 5811 TTATTAATTAA TTATTTATT TAAATAAATT ATTAATTAAG AGATATATCA AAAATTAAAG GTTATTATAA
 5881 CTTCAAATTT AACATATAAT TAGAAATAT ATGACATATACT TTCTGCACT CTCTTGTAT AAATCCAGAG
 5951 AAGCTTATTAG TATATTCCTA ATCAAGTCCA AACCTAATGA AGCTTATATA ATTGTTGAA AACTCAATT
 6021 GCATTAGTTT TAAAGAGTCA CCAAATTCAA AGATAATTC AATGCTTCA TTACCACTAT GGAGAAAATA
 6091 TTTCCTTATG TTAAATGAAA TGAAACAAA CATTCAAACCT AATTTGTGCT TATTAACCA AAGACCCATT
 6161 ACTTAGCCAA GAGTTTAAAC AAAAATTAC ATCATTCTGT ATCATTATTTC ATGACTAGAT ATATATGAA
 6231 ATGAA-GGGGAG TTTTATAGA AAATATAATC ATAGTATTC AACATTAACCT CAGGGAAATTIC CTCAAATAA
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 6371 AACTAAAATA AGGATTAGAA GTACCAAACA TGTTAGAAGA ATCACAGTAA AAGATGATGT TGTTCTGT
 6441 GTTCTTCTAA GTTCTTCAAG TCTCCAGTGT CTCCATTAATG TGCAAAGGGAG AGCCATTAAA TTGTTATG
 6511 TTGATCCCTT CAAAAGCTGC ACCAACCTCC CTAAATAAAC ACTCAAAGCA AAAATGACAA AATGCCCTG
 6581 AGGACCTTAT GTGGGTGCTC TGGCGGGTGC GAGCTGCATA CGAAAGGTCT TTGTCCTTIG TGAGGTGAT
 6651 GTTGTGCGGG ATAGCTTGTG GCATGCTTCC GCGCGGTCA CGCACATGTG CACAGGTGAT GCATGGTGT
 6721 TGGCTTCTTG AGTTTGTAGC CTCGGATGCT TAGTCACCTT GGCCCAATTG GAGTCCAATC AGCTTATAAC
 6791 CCATTCTTCT TCAAGTTATC TTCAAGTTAA GCCCAATTG GCTCTCCAA ATCATCCATA ACTTCACAGA
 6861 ATGGCCCGTT CATCTTAATC CCGGATGCACT AATTATCTC CGGTCTTCAT TTAAGCAAG ATACCACTT
 6931 CTTCATGCTT CATCCATCAA TAGTACACTT CATGTTATCAT CTCTACTAG TATTAGTCG ACAAATCCTT
 7001 GTTGTCTCC AAATTAAATT ATCTCATTTA GTTCCCCTT CGCTTACTTT CCTTAAATTG TGGAAATTAG
 7071 CTCAAGAGAAA TATTAAGTAC CGCAAATGGT CATAAAATTAA ACAAAAAGGA AAATGCAATGA AGATTAAC
 7141 AATGATGAAC GAAATATGCT AAAATGACT ATAAATGAA GTAAATTTAA TGAAATTATC GCACCTCCGAC
 7211 CACCCATTATG GCTTGTAGTC CACCCACCT CTATTCCTTG TACCAATATG GATGGAAAC ATCATTAATT
 7281 AAGCCAAAAA GCTAACATAT AAGGGTTAG TGACAAAGGT AAGTACTAA GTGAAAATA ATCCATTTTT
 7351 CTGTTTTTA CACAACACAC ACATAGGGC AGACCTAGGA TTTCAAAGTA CAGATTGTTG GTGGCACATA
 7421 AGTGTGTGCT GTGACATT TTCTTCTTT TTCTGTTG TGCAACACTG AGGAAAAACG AAAAATTGCA
 7491 AATTCTTATC AATTGTCCTT AAAAATGACA GGGGTGTTG GTGCCACTAT GGACAACAAA GTTGAACITG
 7561 CCTACCGCGG CACACACACA CACACACATA GAGAGAGAGA GAGAGAGAGA AAGAAAGAAA
 7631 GAGAGAGAGA GTTGGGATG TGATACCTT TTAGGAAAA TGAGTTATA TCTTTGATAT TGTTTTT
 7701 TAATGTAATT TATTTATTTA ATCATTATG TTATTAAGTT NTATTTATIN GGTTATGAAA AAAAAAGCT
 7771 TTATACATT GGATTTAACCA TAAACATCA ACAATTAA TCAAAAGAC CAAACATGTG GACAATTATG
 7841 TATATAATT ATTCAACATA GTCTTCTGAGA ATAGTATTAT ATATATAATT AATTCCTCAAT GGTCTTGG
 7911 ATAGTAAGTT CTTATTTTC AAACCTTTCG CAACTTCTT TGCTTACTTT GACACTTTTC TTCCCTAACT
 7981 TTACATATAT ATATATATTA AAGGGCAAAG GTCACTAGGA TATAATATTT TCTTATTATC TACGTTTGC
 8051 CACAAAAGTT TGAACACITGCCACTTTT GTCCCTCCCT AACCTTTCA ATGTTTGGCG ACAAAGTT
 8121 CAAACATTG CGACTTTGAT CATTCTCAA CTTTACCGG CATTAGTTG TGGAGTTGGC AGTTTGGC
 8191 CCTCTTAACTT CGATATTCTC TACTGCTAGC CAAAGGGT TCCAGAGTTT CACACTTTG CTCCCTGACA

RLG 2B cont.

8261 GTCACCAAAAT GTGAGATGTC AAATTTTGC CACATTAGTT TGTGGAGTTG TCCCTTTGG TCCCCCACA
 8331 TTGATATTTC TACTATACGA TCCTTATTTT CTCAAATAAC AACACGTATA TTTCATC:CT AATTGGAAA
 8401 AGAGTTTTAA AA:AAATAAC GACTAGG::: G:GC:GAGTT TTTTTT:ACA AGTTTGATC AAATCATATC
 8471 AAAATTAAAG GTGGAACCGGT GACCACATTA ACCAGAATG TAATTATTC TTGATTTTG ATAATTTTTA
 8541 ATATTTTGTG TGATCTATG TATTTAAAG TAACACAAAG AGAACATAAT CAAAACCCCT AAATGCAAG
 8611 TCTCGCCAA TTTCTCTATC ACTAGTCCTC ACTTACCGATG GCGTTACGTC GCTCTCTCAC TGCTTACAC
 8681 CCTTGTGTC TACTCATTAC AATAACGAA AGTTGAATAT CCATATATTT ATTGGATGT GGAATTGAAC
 8751 GAATCTCGTC AAAATTGAA TTTTGTGAT GGATTTGAGT AGAAGTTGG GCAGAACCGG AAATGATGGTC
 8821 TCGAACGGT TATAAACCTG ATTCTGAGTT ATTACATG ATGTAACCTC TTACAAACGA CCAAGGTTTC
 8891 TTCCAGGTAC CATTGATCT TTTAGAATG TAGTTTCTG AAACACCTG ATTGGATCA AATATCACCA
 8961 ACAACTCTTA AAAACTTGTG TAATCAATG TTTCTCTCAT CTIGATAACA AGTGGAAATGA TTTCTACTT
 9031 AGATTAACCTT GAAAAAAAG GTCCATGTGTC GTCTGGTGG A TCTGGTAAAT GAAGATGGAA GGGAGAGCTG
 9101 ACCTTTAAAGA CACAAACACG TCACCATATC TCTTATTTA TTAAATTTT GCTTGTGGT TATTTCTTT
 9171 TTTCTTTCTT CTTCATCTC TGATCTCCAG ATGGTATGTC GTGTGGATAA TTACACCTA GAGATTGGGA
 9241 ACGATGGAA GGGGTCTGTTG ATTATGCGT GGCGAGTTT TACTTTAACTC AATATTCACCA ACCTAAATTG
 9311 TGATTCCTGTTG TTGAAAATAA GTGCACTT TTTTTGTA TTATCTGTT GCATAGGATC TTAGCATCT
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 9451 TGTTTCTGAA AAAAATTAAC CCTAAAATAA AAATAACCAT TTCAATCTC AAAATTATAA GAGAGAATTG
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 9591 GCAGATTGAA ATTCTATCAG AGAAAGAGAC ATTACAGAA GCCACTGGCA GTATTCAAA TCTTGTATTTC
 9661 CCATCCGTGTC TCATGCACTC TTTTCATAAC CTCCGTGTC TTACATTGGA TAATTATGAA GGAGTGGAGG
 9731 TGGTATTTGAA GATAGAGAGT GAGAGTCAC CATGTAGAGA ATTGGAAAC ACTCGCAAAACCA
 9801 CCCTATTATAA CTTCCCTACCC TCCAGGATTT GTATCTAAGG AATATGGACA ACACGAGTC TGTTGGAAG
 9871 TGCAGCAACT GGAATAATT CTTCACTCTC CCAAAACAC AATCAGAAC CCAATTCCAC AACCTCACAA
 9941 CCATAAATAT TCTTAAATGCA AAAAGCATTA AGTACTGTT TTGCGCTCTC ATGGCAGAAC TCTTTCCTAA
 10011 CCTAAAGGAT ATCCGGATAA GTGAGTGTGAA TGGTATTTAA GAAGTGTGTTT CAAACAGAGA TGATGAGGAT
 10081 GGAAGATCAA CTACATTTAC ATCTACCCAC ACAACCCACA CTTTGTGTC TACTCTCACTC
 10151 TAAGTTTCTG GGAGAATCTG AAGTGTATTG GTGGAAGTGG TGCAAGGAT GAGGGGAGCA ATGAAATATC
 10221 TTTCATAAT ACCACTGCAA CTACTGCTGT TCTTGTCAA TTGAAAGTAT GCTTGTACA TATTCATTA
 10291 TTATTTTAT TTCTTTTTT ATTGCAATA TTCTTATAAT AATACATTTT ATACCACTA TACTAAGATA
 10361 ATAATTACCT AGAGGGATGG ATGCTATGAC ACAGCTGCTA CACTCAGAA ACTCTARTAA GGGCAGTTAT
 10431 GGAAGTCAA TAAATGATA ATGGCATTT TTGATGGTA ATATGGCAA TTAAAGTTT ATTCTGTAA
 10501 AAGCAGTATT TACCAAGTAC TGGCCAGTAG GAGAGGAGAA TATCACCTTT TGCAAAATC TGGTCATGTT
 10571 ACCCAGAATT TAGTTAAATG TAACATTAA GATATTAGGG GTTATCAGGT GACAGATATT GTAGAATAGA
 10641 ACAATATGTA ATATTCACCA AAACTTTT TTCTAAGGTT GTCTCTGTTA ATATGTGCTT TCTTGATTTC
 10711 ATTGAAATTG CATTCTTATGTT TTTTAGGTTG TAAAGTGTATT GTCTCTCAA TAAATCCGA AATTTTTAA
 10781 TTAAAAAAAG AAAAACAAA AGTAAATTGTT TGATATGGAG AGCAGTGGTA TCATTTAGTA TATTTAAAC
 10851 AGATTTGAA TTAGTTCT TATATAAAAG CTGTGTATAT AGTTAATTG GTTTTACATC ATTTTTCCAT
 10921 GTGGTGTGTC AGTGTGTGAA AGCAGGTGGT GTTTCTGGA GCTTATGCCA ATACGCTAGA GAGATAAAA
 10991 TAGGCACACTG CCATGCATTG TCAAGTGTGA TTCCATGTGA TGCAGCAGTA CAAATGCCAGA AAGCTT

SEQ ID No: 23

RLG 2 B a.m.

MSDPTGIAGAIINPIAQTAALPVTDHGYMISCRKYVRVMQMKMTELNTSRISVEEHISRNRNHLQIP
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GAVI?EKTDPFIAQEAIADYLGQLNEKTTPARADKLREWFKKNSDGGKTKFLIVLDDWQLVDLEDIGL
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CGLPIAIKTMAC?LRNKRKDAWKDALSRIEHYDIHNVAPKVETSYHNLQEEETKSTFLMCGLFPEDFDI
PTEELMRYGWGLKLFDRVYTIREARTRLNTCIELVQTNLLIESDDVGCVKMHDLVRAFVLGMFSEVEH
ASIVNHGNMPGWPDENDMIVHSCKRISLTCKGMIEIPVDLKFPKLTLIKLMHGDKSLRFPQDFYEGMEKL
HVSIYDKMKYPPLLAPRCSTNIRVLHLTECSLKMFDCCSIGNLSNLEVLSFANSHIEWLPSTVRNLKKL
RILDLRFCGDLRIEQGVLKSFVKLEEFYIGDASGFIDDNCNEMAERSYNLSALEFAFFNNKAEVKNMSFE
NLERFKISVGCSFDENINMSSHSYENMLQLVTNKGDVLDLSKLNGLFLKTEVLFLSVHGMNDLEDVEVKS
THPTQSSFCNLKVLIISKCVELRYLFKUNLANTLSRLEHLEVCECENMEELIHTGIGGCGEETITFPKLKF
LSLSQLPKLSSLCHNVNIIGLPHLVDLILKGIPGFTVIYPQNKLRTSSLKEGVVIPKLETLQIDDMENLEE
IWPCESGGGEKVKLRAIKVSSCDKLVNLFPRNPMSSLHHLEELTVENCGSIESLFNIIDLCVGAIGEEDN
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GNHESEEQVTLSISLS

SEQ ID NO: 24

SEARCH NO.:

		→ 2.5	
RLG2A	-- CGAGAC-ACATCTCA-ACGCTTGTGAGAATGTTGGAGAAAGAATGTTTAATATATTTGAGCCCGTTATAGGCCAAAGCACGCC	10	100
RLG2B	-- TGGCGAAAGCTAACATCACTAAAGCTGAGAATGTTGGAGAAAGAATGTTGGAGAAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	20	100
RLG2C	-- AACAC-ACGCTTGTGAGAATGTTGGAGAAAGAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	30	100
RLG2D	-- -ANGCA-CTTGCTCAAAGTGTGAGAATGTTGGAGAAAGAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	40	100
RLG2E	-- -ANGCA-CTTGCTCAAAGTGTGAGAATGTTGGAGAAAGAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	50	100
RLG2F	-- CTGGAGAC-ACATGTCATGAACTGATGAAAGGTGAAATGGTGTGAGAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	60	100
RLG2G	-- CGAGAC-ACATGTCATGAACTGATGAAAGGTGAAATGGTGTGAGAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	70	100
RLG2H	-- TTAGGGCTTGTGAGGAGCTTGTGAGAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	80	100
RLG2I	-- AT-GCA- --GAGAGCTTGTGAGGAGCTTGTGAGAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	90	100
RLG2J	-- GAAAGA- --GTCGTC-ACMAGAAAGAACCTGCAATTTGAGAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	100	100
RLG2K	-- TTTGAGAC-ACCATGTCATGAACTGATGAAAGGTGAAATGGTGTGAGAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	110	100
RLG2L	-- G- --GCTGAGAGG-CTGCTGAAGAAAGAAATTTGTTTAAATTATTTGTTGGCCAGTTATAGGCCAAAGCACGCC	120	100
RLG2M	-- CATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	130	100
RLG2A	CATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	140	100
RLG2B	CATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	150	100
RLG2C	CATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	160	100
RLG2D	CATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	170	100
RLG2E	CATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	180	100
RLG2F	CATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	190	100
RLG2G	TATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	200	100
RLG2H	TATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	210	100
RLG2I	TATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	220	100
RLG2J	TATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	230	100
RLG2K	TATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	240	100
RLG2L	TATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	250	100
RLG2M	CATTGCTTATCGCAAGCTGTAGCAATGTTGGAGAAATGTTGGAGCCCGTTATAGGCCAAAGCACGCC	260	100

<u>AAGAAATTTCATTAACNTGCAAGCGTATGCTGTTGACTGAACTTCAGTTGACCTAACCTTACGGATTTGAACCTTATGCTAAGCTTATGCTAAGCTTATGCTAAGCTTATGCT</u>	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2A	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2B	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2C	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2D	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2E	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2F	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2G	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2H	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2I	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2J	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2K	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2L	1010	1010	1040	1050	1050	1070	1080	1090	1100
RLG2M	1010	1010	1040	1050	1050	1070	1080	1090	1100
<u>AAGAAATTTCATTAACNTGCAAGCGTATGCTGTTGACTGAACTTCAGTTGACCTAACCTTACGGATTTGAACCTTATGCTAAGCTTATGCTAAGCTTATGCTAAGCTTATGCT</u>	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2A	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2B	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2C	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2D	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2E	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2F	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2G	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2H	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2I	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2J	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2K	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2L	1110	1120	1130	1140	1150	1160	1170	1180	1190
RLG2M	1110	1120	1130	1140	1150	1160	1170	1180	1190
<u>AAGAAATTTCATTAACNTGCAAGCGTATGCTGTTGACTGAACTTCAGTTGACCTAACCTTACGGATTTGAACCTTATGCTAAGCTTATGCTAAGCTTATGCTAAGCTTATGCT</u>	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2A	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2B	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2C	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2D	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2E	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2F	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2G	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2H	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2I	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2J	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2K	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2L	1170	1180	1190	1200	1210	1220	1230	1240	1250
RLG2M	1170	1180	1190	1200	1210	1220	1230	1240	1250

SEQ ID NO:

GETT-----LGEVWKEFQHIVAVIGENTDPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 40
 RILG2A protein in CTKMHLRKKVKERKFIIAEVGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 98 - 4/
 RILG2B protein in QKTHMLRKKVKERKFIIAEVGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 100 - 4/
 RILG2C protein in NTRK - AKAEVAKRKKVKERKFIIAEVGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 98 - 43
 RILG2D protein in EVAK - RK-----RKEVAKRKKVKERKFIIAEVGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 90 - 44
 RILG2E protein in GRND - AKVEEVAKRKKVKERKFIIAEVGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 100 - 4/
 RILG2F protein in DEDTMQRKLRKKVKERKFIIAEVGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 97 - 47
 RILG2G protein in GRUDD - EELKVEGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 89 - 4/
 RILG2H protein in CTKS - -KEVKGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 94 - 49
 RILG2I protein in ERGR-----GDKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 100 - 5/
 RILG2J protein in LEDTMKHLRKKVKERKFIIAEVGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 82 - 5/
 RILG2L protein in AEE-----FSYAVLVEVIGKTFPIAICQAVDGLIEKESTPARADKLEMFTKREMFYLTLDWVQSOVLEIDGLSPFPNG 92 - 53

VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2A protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2B protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2C protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2D protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2E protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2F protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2G protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2H protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2J protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2K protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2L protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD
UJLG2M protein VDFKVLTLSRSQVCTMVEANSIANGLLIEAJSLSLQFQFVETS-E---PELKNGEDIVRKCCGPIAIKTMACTLRNKDQWADLSRLEHD

:S--VAPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL
 RLG2A protein IIN--VAPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 294
 RLG2B protein IIN--VAPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 293
 RLG2C protein IIN--VAPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 291
 RLG2D protein IIN--VAPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 283
 RLG2E protein IIS--VVPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 291
 RLG2F protein IIS--VVPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 293
 RLG2G protein IIS--VVPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 297
 RLG2H protein IIS--VVPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 289
 RLG2I protein IGS--VSEVREVKFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 294
 RLG2J protein IGS--VSEVREVKFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 289
 RLG2K protein IET--VAPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 293
 RLG2L protein IXX--VAPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 275
 RLG2M protein IIN--VAPKVFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL 285

:S--VSEVREVKFESTYLNQDEEKTSTFIMCGLEPPDFDPIPEELRYGAGLKLFDVVTYIREARNRNLTCERLQVQHILLESDDOVCVQHDLVRATVL
 GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS
 RLG2A protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 390
 RLG2B protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 391
 RLG2C protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 389
 RLG2D protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 381
 RLG2E protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 387
 RLG2F protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 389
 RLG2G protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 394
 RLG2H protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 386
 RLG2I protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 391
 RLG2J protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 386
 RLG2K protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 392
 RLG2L protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 371
 RLG2M protein GMF SEVEN-ASPARTHEN--MPCGEND-IVHSCKRISLTCKGSEEPFDLAKPENPTILAKMHDCKSLRPDPFTEGMELQVLSYDCKYPLPSPPCS 383

	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -				
RLG2A protein	VNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	410	410	410	410
RLG2B protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	450	450	450	450
RLG2C protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	470	470	470	470
RLG2D protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	480	480	480	480
RLG2E protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	470	470	470	470
RLG2F protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	470	470	470	470
RLG2G protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	470	470	470	470
RLG2H protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	480	480	480	480
RLG2I protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	470	470	470	470
RLG2J protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	470	470	470	470
RLG2K protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	465	465	465	465
RLG2L protein	TNRLVHLHFC SURFDCCSISGILNLLEVLSFANSISIEMLPSTIGNATKRLDUNTCYLRGTRTGNQG - P0 -	480	480	480	480

SEGMENTO:

SEQ ID NO:

	810	810
AC15-2A	TAAGTACTTGTTTACCCCTAACGG - 66	779
AC15-2B	TAAGTACTTGTTTACCCCTAACGG - 67	777
AC15-2C	TAAGTACTTGTTTACCCCTAACGG - 58	777
AC15-2D	TAAGTACTTGTTTACCCCTAACGG - 57	778
AC15-2E	TAAGTACTTGTTTACCCCTAACGG - 60	721
AC15-2F	TAAGTACTTGTTTACCCCTAACGG - 61	781
AC15-2H	TAAGTACTTGTTTACCCCTAACGG - 62	738
AC15-2I	TAAGTACTTGTTTACCCCTAACGG - 63	722
AC15-2J	TAAGTACTTGTTTACCCCTCAC - 64	784
AC15-2L	TAAGTACTTGTTTACCCCTCAC - 65	699
AC15-2N	TAAGTACTTGTTTACCCCTAACGG - 66	778
AC15-2O	TAAGTACTTGTTTACCCCTAACGG - 67	763

111

SEQ ID NO:68

RLG3 (real RLG3)

[Strand]

1 AATGCCAAA GAACTCGGAG CAAGAGCTAA GTTAGAGCAT CTATTGAGC TCATTATCAT GGTAGATGTC
71 ACTCAAGCAC CCAACAAGAA CACAACTCAA AGTAGTATT CAGAACAGTT GGATTAAGAA CTGCAAGAAG
141 AGACCTTGTG CGTAAGAGCA GCTAGGGTAA GTGGGAGGTT AAAAATGCTT ACAAGGGTGC TGGTGATATT
211 AGACGATATA TGGTCAAGGC TTGACATGGA GGAACCTGGG ATTCCCTTG GATCAGATAG ACAACACCAC
281 GGCCTGCAAAA TCTGTGAC TTCAAGAAGT ATTAGTGCTT GTAAACCATG GAGAGCTGAT AGAATCTTTA
351 AAATACGAGA AATGCCACTG AATGAAGGCAT GGCTTTTTT CGAAAGAACCA GCTAAAAAAAG CTCCGAATCT
421 GCATCAAGTA GCAAGAGATA TCGTGGAGGA GTGTGGTGGG C

RLG4
SEQ ID NO:69

1 GAATTOGGTG TTGGTAAGAC AACTCTTGGCC TCTTCGTGTT ATGATGAAAT CTCTAGCAAG TTTGATGGTT
71 GCTGCCTTCT AAAAATATCT GGAGGAATC AAGTAATAAA GACCGTATAAC AAAGATTGCA AGAAAAAATC
141 ATTTTGATG TTTTGAAACA AGAGCAAGTG GGCGTAGGGA GAGTTGAAAGA AGGAAAGGCC ATGATAAAGG
211 ATAGGTTPACA ACATAGAAAG GTATTGATTG TGCTTGATGA TGTCGACAAC GTTGAGCRGC TAGCTAGAAC
281 AGTTGGCTG ATCACATGAT TGCTTGCTG AACGTACCCC CATAATAATC ACAACTAGAG ATGAAACATGT
351 ATTAATIGCA CACAAAGTAG ATCTGATACA CAATATAAGC TTGTTAACAC ACGATGAAGC TATGCATCTC
421 TTCTGCAAGC AACCAACACG CGGTCAACAAA CGTATACAAG ATTATGAGCA ACTTTAAAAA CATGTGGTT
491 CTTATGCTGG TGGGCTTCCA CTAGCACTGT CGAC

SEQ.ID NO: 70
 RLGI-E169
 [Strand]

1 ATCGTAACCG TTGTTACCG AGCCGTCCTCCTCCATC TTTTGTCTATA TGTCATATTC TCATINNATTN
 71 TGCACACATT AATTTCGGG TTATTTAA TTAATTTTA TTCCACATGT CATTTTATGA GTTTTCTAT
 141 TTATTCAGT TTACATAAT ATTAACTATG AATACAACTA AATCCATATG TATTTCCTT TAATAAACG
 211 CATAATAAT ATAGATTTAA ATCATATAAT ACATAGGTT AACTCATATA ATACATATGT TCATCCCCAG
 281 TTATTTATA TGTCATCCTC TTATTTATT TATTATTTAT TTATTAGAGT AGATGATCTT TGATGATATA
 351 AAAATTAAAT TTGTTCAAAA TTAAATATTA TTAAATATGCA CACAATTGTA ATAAATTTAA AAAAATGGN
 421 CCCACCAATTG TTCCATCACT TTTCAGCTC ATCAATATCG TGAGTATTCT CCTTCGTTTC CACCTTAATC
 491 AATATTTCCA CGGAATGACA GACTCTTAG CGGTTCTGTA ATTTCGGTT CGACACTGTT CATTGAGGA
 561 GATAATAATG CAAATGGAGC TGCTCCTAG TGCTCTGAG ATGAAAGGTG ATGTTATGT GAAGANAATG
 631 TCACCGATCCTC ATCTCCATCC GGAACCCACC ACATPATCAG TGTCACCA AACCACCAA AACGGYGGAA
 701 GTAGGRKAC WRKAAGTCGA TGAAGAATAG ATTATTTTG TCCCTCATGGG CTGACTGAGG AGCGGGTTTA
 771 GTTCATCATT TTCTTCTTGA CAAGAATTG TGCGTCCATC GAATTTCATC ATCGACAAAG AAGTTTCATC
 841 TCGCAATGTT TTGTTAAACA ATTTCATTC TTTTATCTT TTGCTGTGAA CCTCTCAATT SCAACTTGA
 911 ACTTGCAACT TTGGGGCCCA CAAATTGTTG GTGGGGCTTA ATTAAATCCA CATAATTCATC GTPAACAAATA
 981 ATTCAATTCG ATCTCTGTT ATCCCAATTC TCAACATCTC TTGATAATG AAATCAATTC GCGCTTCATCC
 1051 ATTTCATCCTA CATCTATATC ATTATCTCTG CTCTTATCAT ATTAAACCGT GGCTGAAATC GTTCTTCTG
 1121 CCTCTTCTGAG AGTGGTGTGTT GAAAGCTGG CATYTGAAAGC CTTGAAAGAAG ATGTTTGGCT CAAAAGAAT
 1191 TGATCTGAG CTAAAGAATG TGAAGGAGAC ATTAGACCAA ATCAAGATC TGCTTAACCGA TGCTTCCCCAG
 1261 AAGGAAGTGA CTAATGAAAGC CGTAAAGAAG TGCTGTAATG ATCTCCAAAC TTTGGCTTAT GACATAGACG
 1331 ACCTACTGAG TGATTTGCA ACTGAAGCTG TTCAWCGIGA GTTGACCGAG GAGGGTGGAG CCTCCCTCCAG
 1401 TATGTTAAGA AAACAAATTC CAAGTTGAGC CACAGTTTC TCACAAAGTA ATAGGATGCA TGCAAGITTA
 1471 GATGATATTG CCACCAAGGTG ACAAGAAGCTG TGAGGGCAA AAATAATATCT TGGTTTAAGT GTGATAACAT
 1541 ATGAAAAGCC AAAATTGAA AGGTATGAGG CGTCTTGTG AGATGAAAGC GTGACTGTCG GACTGIGAAGA
 1611 TGATAAGAAA AAATTCCTGG AGAACGCTGG CGGGGATAAA GATGAAATCAG GGAGTCAAA CTTCAGCATC
 1681 GTGCCCATAG TTGGTATGGG TTGAGTTGGT AAAACAATC TAGCTAGACT TTGTTATGAT GAAAGAAAG
 1751 TGAAGGATCA CTTCGAACTC AGGGCTGGG TTGTTGTTTC TGATGAGTTG AGTGTGCTCA ATATAAGGAG
 1821 AGTTATTTAT CAATCTGAG CGGGGAAAAA GAAGGAGTTG GAAGACTTAA ATCTGCTCA AGAACCTCTT
 1891 AAAGAGAACCT TTAGGAACCA CCTATTCTA ATAGTTTGG ATGATGTTG TGCTGAAAGC TATGGTGATT
 1961 GGGGAAAAATT AGTGGGCCCA TTCCCTGCCG GGTCCTCTGG AGATGAAATA ATCATGACAA CTGGAGAGGA
 2031 CCATTCCTCAG AGAAAGCTGG CCTTCTCTCA TGACACCCCT CTGAGGGTC TATCACAAGA TGATGCTTTC
 2101 TCTTGTGTTG CTCAACACCC ATTTCGGTGA CCAAACATTG ATTCACTATC AACACTAAGG CCACATGGAG
 2171 AACTTGTTGTTG AGAAATTGATG ATGGCTTAC CTCTAGCYTT AAGAACACTT GGAGGTTTATG TAAGGACAAA
 2241 AACAGACGGAG GAAACATGGAG AGGAGCTGGT GGATGAGTG ATATGGAGGT TAGGAAGAG CGATGAGATT
 2311 GTTCCGGCTC TTAGACTAAG CTACAAATGAT CTTCCTGCCW CTITGAAGGT RTTRTTGCA TAYTGCTCT
 2381 TGTTCCTTCAAGA GAACTATGAG TTGACAAGG AGGAGTTGAT CTATTTGTGG ATGGCGAGAG GCTTCTTICCA
 2451 CCAACCAACT AYAAACAAGT CAACCAAGC KTGGGTCTT GAATTTCATR AAGAGTTRTT GTCAAGRTRCR
 2521 TTTTTCAAC AGTCCTCAA TRCAACATC TTGTTGIGA TGCAATGACCT ATGAAATGAT TTGGCTCAT
 2591 TTGTTGCTGG AGAATTTTT TCAAGGTTAG ACATAGAGAT GAGGAAGGAA TTGAGGATGS ATCTTTGGA
 2661 RAAGCACCGT CATACTGAG TTGTTGIGA GRATACATCA GGTTAGCTAA RGTTGGAGCC ATTTAGAGGA
 2731 CCTAAAAAATT TGAGAACATT TTGAGCTATG TGTTGTTGGG TGTTAGAAGA TTGGAAGATG TTCTACTTAT
 2801 CAAACAAGGT CTGATGAC WTACTTCARG ATTACCAT TGTAAAGGTC CTRAKTTGCA TYTRRTCTTAY
 2871 ATAASYRAG STACCARAA TGTCGCGTAG TAGTAAASCAC TGTCGGTATC TTATATCTP WGRAACTTWA
 2941 ATCACMCATT TACCGGAAWA TTCTCTGCAAT CTTATTAAT TACARACCCCT GATTTGTTCTT GGCTGTGAMP
 3011 ATTATGTTAA KITGCCCAAR ACCTCTCTAA ASCTTTAAAAA TTGAGCATC TTGAGCATG GGGTACTCC
 3081 KAAKTTAAR CAACTGCTTCT TARGGATTG TGARTTGAAA ARCTCTACAA CTCTCTTYMG TAACATTGGC
 3151 ATACCAATTA CGGAGCTTAA GAACCTGCAAM AAYCTCCTAAG GGAACATTG TATGGGGG CTGGGAAAAA
 3221 TGGAAAAATCC NCTGGAATGC ACCTTAAGGC AACTCTCTC A: AAAAAAGGT TWAATGARTT ANAAACTGGR
 3291 WTKGGGGCTT ATRAATTAA TTGTTCTCGA ATGGGAACA CTTGAAAAAA NAAGGTCTC AATGAAATTGA
 3361 ATCCCTCATA ATGGTAYTCY AMWAARRRY YWTARWAT IWMGKAWRK GKTTTATRR TKTMYRAW
 3431 WAGRTKTR KARSTAGGTGTT TCATCCAACT ACCCAAGTGG GAAATAGAT GATATTTCA GGGCYTACTG
 3501 ATGAGATGTTG GAGGGTATG ATAGGGTNTC TTGGGGCGGT AGAGAAATA AGCATCCATT TTGTTAATGA
 3571 AATAAGATAT TTGTTGGAAAT CAGAAGCAGA GCGCAACTAAG TTCTTATGAA ATTAAAGAA TTGGGATTAA
 3641 GTGGAATGTT AAAATTGGT GAGTTTGGG GAGAAAAAGG AGGATAATCA TAATTTAAT ATGGGAGGA
 3711 GCCTACACCT TTGTTAGGAG TTGATGTTGAG CAGCTTGGGAG CAGCTTGGGAG CTTGAGGT GTCCAGATAG
 3781 CATGGAGAAT TTGTTATGCA ACATGTTGAG TTCAATNACA TCCGCTCTCT TCCCAACAGG AGGAGGACAG
 3851 AAGATCAAGT CACTTACCAT CACTGATTGC AAGAAGCTT CGAGAGAGGA GTTGGGAGGA CGAGGAGAGG
 3921 CAAGAGTGT TATAAACCTCA AAAATGCAAGA TGCTTGAATC AGTATGATA CGTAATTGGC CAAATCTGAA
 3991 ATCTATCATTGAA TTGTTGAGTT GCTTCATTC CTCGAACAGA TTATATATAT CAAACTGTCG GAGTRTGGAG
 4061 TCATTTCTG ACCATGAGTT GCGAAATCTC ACCTCTTAA CAGATCGAAC GAGGAGACAC CGATTCTCGT

RLG1-E169
[Strand]

4131 ACGAACGGTT ACGATTCCGAC TGGCCGTGGT TTT

SEQ ID NO: 70

Further Characterization of RG2 Family Members:

Further sequencing of cloned RG2 polynucleotide sequences, as discussed above, identified additional RG2 species, listed below. Additionally, further sequencing of the 5' sections of RG2 sequences listed above resulted in modified and/or new sequence 5 information, also listed below. The AC15 sequences found in the 3' sections of RG2 family have not changed.

Listed below are: four full length species, RG2A, RG2B, RG2C and RG2S; two near complete, but with a gap in the largest intron, RG2D and RG2J; three nearly 10 complete RG2 gene sequences, RG2K, RG2N, and RG2O. The deduced translation products (polypeptides) encoded by these RG2 species are listed below. The polynucleotide sequences do not contain any gaps (as with some of the polynucleotide sequences), because all of the gaps in the sequences are in introns, *i.e.*, there are no gaps in exon, or coding, sequences.

They include: an RG2A polynucleotide sequence (SEQ ID NO:87) and its 15 deduced polypeptide sequence (SEQ ID NO:88); an RG2B polynucleotide sequence (SEQ ID NO:89) and its deduced polypeptide sequence (SEQ ID NO:90); an RG2C polynucleotide sequence (SEQ ID NO:91) and its deduced polypeptide sequence (SEQ ID NO:92); an RG2D polynucleotide sequence (SEQ ID NO:93) and (SEQ ID NO:94), and its 20 deduced polypeptide sequence (SEQ ID NO:95); an RG2E polynucleotide sequence (SEQ ID NO:96) and its deduced polypeptide sequence (SEQ ID NO:97); an RG2F polynucleotide sequence (SEQ ID NO:98) and its deduced polypeptide sequence (SEQ ID NO:99); an RG2G polynucleotide sequence (SEQ ID NO:100) and its deduced polypeptide 25 sequence (SEQ ID NO:101); an RG2H polynucleotide sequence (SEQ ID NO:102) and its deduced polypeptide sequence (SEQ ID NO:103); an RG2I polynucleotide sequence (SEQ ID NO:104) and its deduced polypeptide sequence (SEQ ID NO:105); an RG2J polynucleotide sequence (SEQ ID NO:106) and (SEQ ID NO:107), and its deduced 30 polypeptide sequence (SEQ ID NO:108); an RG2K polynucleotide sequence (SEQ ID NO:109) and (SEQ ID NO:110), and its deduced polypeptide sequence (SEQ ID NO:111); an RG2L polynucleotide sequence (SEQ ID NO:112) and its deduced polypeptide sequence (SEQ ID NO:113); an RG2M polynucleotide sequence (SEQ ID NO:114) and its deduced 35 polypeptide sequence (SEQ ID NO:115); an RG2N polynucleotide sequence (SEQ ID NO:116) and its deduced polypeptide sequence (SEQ ID NO:117); an RG2O

polynucleotide sequence (SEQ ID NO:118) and its deduced polypeptide sequence (SEQ ID NO:119); an RG2P polynucleotide sequence (SEQ ID NO:120) and its deduced polypeptide sequence (SEQ ID NO:121); an RG2Q polynucleotide sequence (SEQ ID NO:122) and its deduced polypeptide sequence (SEQ ID NO:123); RG2S polynucleotide 5 sequence (SEQ ID NO:124) and its deduced polypeptide sequence (SEQ ID NO:125); an RG2T polynucleotide sequence (SEQ ID NO:126) and its deduced polypeptide sequence (SEQ ID NO:127); an RG2U polynucleotide sequence (SEQ ID NO:128) and its deduced polypeptide sequence (SEQ ID NO:129); and RG2V polynucleotide sequence (SEQ ID NO:130) and its deduced polypeptide sequence (SEQ ID NO:131); and, an RG2W 10 polynucleotide sequence (SEQ ID NO:132) and its deduced polypeptide sequence (SEQ ID NO:133).

Characterization of New RG Family Groups and RG Species:

Further BAC insert characterization and sequencing, as discussed above, 15 identified new RG polynucleotide sequences. The new sequences were characterized as belonging to new RG families; designated RG5 and RG7. These RG polynucleotides sequences, and their predicted translation products (the polypeptides which are encoded by these sequences) are summarized and listed below.

Identified and listed below is an RG5 family member, designated as the RG5 20 polynucleotide sequence set forth in SEQ ID NO:134, and its deduced polypeptide sequence (SEQ ID NO:135). This sequence contains an NBS region sequence.

Also identified and listed below is an RG7 family member, designated as the RG7 25 polynucleotide sequence set forth in SEQ ID NO:136. No deduced polypeptide sequence is given for the new RG7 family member as this sequence appears to be a pseudogene.

RG2A polynucleotide sequence (SEQ ID NO:87)

AAAAGTTCATATCCAAGCTTGCCCTCCAACCTCTAGCTCCTTCAATGGCACC
TCCTTCTCTTCAAAGCACACAAGAACACTTTCAAGCTCAACCACACTCA
30 CACAAAGCTCTAGAACGAGGGTTAGGGCACATTAGGGTTTGCTCTCTGG
AAATGGGTCTAAAGTGAGGCCATAATGTTCCCTATATAAGGCTCACTC
CCACAATTAGGCTTCAATCTGAACGTANTACGCCAGTGTACACTATGG
TACGCCAACGTACTCGGTAGTCTCCCGGTCAANAATACACTCATGAGTA

CGCGCAACGTACTTCCCTTACGCCAGCGTACTCAAAGCAAACATT
TTTCAGGACTAATTTGACAACTTGAGGAAAGAAAAGGATCAAAGANA
TATACTTGAACTCCGGGATGTTACAATGAAGTTGANACCTGGCTAAAAA
ATTAAATTGGTGTGGAAGCCGTTGGCTGAGCAAGCAACAAGGGTAAAAT
5 TCGTAATCTACAAATGGTGTATTTCTATTCCTTATTATTTACTT
GATTACGGTAGTTTTCTTACAAAAAATATTAAAGTTGATAAAG
TATGCCACTAAAATTGACTTTCCAAAACATAATGTCAAATGGTGCCT
ATATGTATCATGTTGTATTANATAATGAATATGATGATNCTGTTCTATT
AANCCGAAAAAATTATCTAACGTTATTTATTTGGAAAACAAAGTTGTGAT
10 TTTNGCATAATATAATCAAATCCNCTTGTNTGGGAGGTGGATAAATG
TGGTAAATTANAACAAGTGTTCACNTGAAGGGNTGGAAAGGTTGA
AAAAAGTTAAAATGATAAAATGTTACACAAATGTTGATCCGACTGAAT
ATNATGTTAAGGATNATTGTTAAATTGTTGATATAGTAAGCATAA
ATATTAGAATTGACTAAATTATAAGTTATNCNAACTGGATTGAAA
15 CATTGGATATANATTAGGAATGAAAATGAGCAACCCTAACATACTTAT
CTTGGTAGTTGGTTATTATTTTATTANAATATAGAACATCCCTT
TATTTAAACCCATATTGTGGACGGACTTGAATAAATGGGAAAAATGTAC
CTTGCTATTAGCACAAAAAAATTATAAAATGTACATTGCTATTAGCA
CAAACAAAAAAACTATCCTTTGCATTAGGTACAAAGAAATA
20 TAAAATGGGAAATGTGTTGCTATTAAATGCACTAAAGAAACTATTTGC
CTTATTAAACCGGGTAAACCAATAGAAAAATGGAAGTACATTGTCATT
AGCATGAAAAAAATAACTTCCATTGGCATCCGGTCACAATAATAG
AAAAATGAAAGTACGTTGCTATTAGCAGAAACTAACTTCCTTTCTTT
TTGGCATCGTATCATAAAATATAGACTAAACATGTTAGTTACATT
25 TAATACATTGAAATGTCTAACCCACATGTTATTCTATAAAAGGGAAATG
TAATTACTTATTCTTGATTCTTGCTTCTTTAGTACCCAAAACAT
CCCTCTATCCATCTATTCAACTAAATAATGAAAACATATTCTCCA
TTGTAGGGATGTTATAAATTGTAATTGTTTATGCAAAAAAGTGT
TTTGTAACTAGATTAACGAGATTCACTTTAGCATTAGGAGAAGTT
30 CATCCATCTTGATATGAAAGTGCAGCCAAGTTCTTAACATGGAATA
TGAGGTCCCTATATGCTAAAAAATAGCAAATGAGAAATTTTAAATTG
GATCCCCATAAAAGAAAATTGTTAATGGTTTTAATATTGGTCAATG
TGTCCACCGGATGAGCATAACTAGTTATAAGGGTAAAGGTGGTT
GGTGGGCCATTATCTTATTCTAAAGTCAGAATTAAGTAAAAAA
35 AAATTATAAGATAAACCCATAAGGATAAAAAATCATTATTGGACCA
AAGACCAAAGTGTAAAGGGCTGTTGTTTTGTGAAGAGCTGTGC
AACCACTTTGTCTGCCGCACAGACAACGTGCAGACATATGCCCTCGC
AGAGTGTGTTGTTTGAAAGTGCAGACACAAAAACGTCTGCGCAG
GTCATCCTGGCGCATATATGTCAGTCTCAAAGGTCTCAGACCTC
40 ATTTAACCAAAAAAAAGACCACCGTTTTTTTTNTTC
TTCTCTGTAGCTGAAATGCATTAAATCTTATGACATGAAATTAA
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TCTCTTGTGCAGACTGTGGACATTGGCACCTCTACCGCAGAG

ACTTGCAGATGTGGTCCGCAGACTGCAGACATTGGCTCAAATAACA
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AAGAGTTATCTAAATTTAGTTAGTCGGTAGATCAGTCTCACATTAAACCG
5 GGTAAAGTGTATGTGTACCGCGCACCTGAAAGGTTGAANGTAACCT
CCA AACTGAANCAANAATCGATATGAAGTATCAAGTTAGAGGTTCAATTG
GTGAAGGAATCAGCTGGAGGTTGGGAATCGAGCTCCACTATTAAGGTA
AAATCCATAACCCTAAATGTTGGTACGCTCATATATCAAATTGCGTGT
TGTGAATGAAAAAAGCATGCTAAAAAACCAGTGTAAAGGCACGGTATAT
10 GACATATTAGTTACTGATAACAAATTATGATAATTGGGTTACGT
AAGTTAGGATTCTGTAATTCAACCAAATGTAATAGTTTGAGTCTATC
TATGTATTGGGAATCACATTAGCAACGGGATTGTACTAGTAATTGAA
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ACA TCTAATATTAAATAGAATGTATCTGATATTGAATTAAATGCTTAAT
15 GTG AACATAGACCTTCCATTACTAATGCCTAATTATTAGTTCTAAT
CAATAAATTAAATTCTGTTTATGCTTCTAAGACAATAAAATCCATG
ATTACCTTAAATATTAACAAAATGACCATAAAATAAAATTTAG
GAT ACCAAACCCCCCGCCATGCCAATGTCTAAATTCTTGATGCTT
TGCTTCCCTCTTCCCTGTTAGTCTATTATTCTGGAGAGTTGAGAG
20 AGTTTCATACAAGAAAATTCAAGAAGAAAGCAAAGGTCAGGTATTCTC
TTTCTTAATTATGTATTAACCTACAAGCATTTCACGATCCATGGT
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25 GAGATAGAGATATGTTAAAACGGCTAGAAAATTGTTAATTGAAATT
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30 GGAGATACTGTGTACCTCGTCTAAAGTCCTGATATTAAACACCTTC
CAWGTCTTAAGGGGTTAGCTGGGAAGGAGGATTTCTATTCCCATT
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TCTTTATGCTGCAGGGAAAAAGACATCAACTCCTCTATTATAAGATC
35 AAACAACAGGTAAACCAAGATCTTGTGCTNNATAATTCTAACNACA
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GACTCTGATTAATGTGAAGTGAATATTAAAGGTAAATTATTTCATGT
TCCTAGTNGCCTATTAAATTAAAGGCCTTGTAGTCNGGATTTGGATGT
40 ATTCTTCATGATGATGTCAATCTCTAAACCCATTCAATTGTTGGTTG
AATGTTGACTCTATGTCAAGGATGAATTCAAGGGAAAGAATTGTTCATCA
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RG2A deduced polypeptide sequence (SEQ ID NO:88)

MDVVNAILKPVVETLMVPVKHIGYLISCRQYMREMGIKMRGLNATRLGVEEHVN
 RNISNQLEVPAQVRGWFEEVGKINAKVENFPSDVGSFCNLKVRHGVGKRASKIIEDI
 DSVMRREHSIIWNDHSIPLGRIDSTKASTSIPSTDHHDEFQSREQTTEALNALDPNHK
 5 SHMIALWGMGGVGKTTMMHRLKKVKEKKMFNFIEAVVGEKTDPIAIQSAVADY
 LGIELNEKTKPARTEKLRKWFDNSGGKKILVILDDVVWQFVDLNDIGLSPLPNQGV
 DFKVLLTSRDKDVCTEMGAEVNSTFNVKMLIETEAQSLFHQFIEISDDVDPELHNIG
 VNIVRKCGGLPIAIKTMACTLRGKSKDWNALLRLEHYDIENIVNGVFKMSYDNL
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 10 HTNLLMEVDDVRCIKMHDLVRAFVLDMSKVEHASIVNHSNTLEWHADNMHDSC
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 MWNLKEIWPCEFNMSEEVKFREIKVSNCDKLVNLFPHKPISLLHHLEELKVKNCGSI
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 HHNQQQPIILPNLQELILWNMDNMSHWWKCSNWNKFFTLPKQQSESPFHNLTTIKI
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 KNNKSGAGEEGIPRVNNNVIMLSGLKILEISFCGGLEHIFTSALESRLQLQELKITFC
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RG2B polynucleotide sequence (SEQ ID NO:89)

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RG2B deduced polypeptide sequence (SEQ ID NO:90)

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 DEGSNEISFNNTTATTAVLDQFELSEAGGVWSLC**Q**YARE**E**I**V**GCYALSSVIP**C**YAA
 25 GQM**Q**KL

RG2C polynucleotide sequence (SEQ ID NO:91)

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 T**C**T**A**TT**A**AA**A**TT**C**TT**A**AA**A**CT**T**C**T**TT**A**AG**C**C**A**GG**G**T**G**A**A**T**C**A**A**
 TG**C**TAG**A**CC**C**ACT**G**TT**A**TT**C**CAT**G**A**T**AT**G**C**T**G**A**T**C**A**A**TT**G**T**T**GG
 CT**G**C**T**AC**G**AT**G**C**A**GG**G**G**C**T**A**CC**A**AG**A**AT**A**T**G**G**C**C**A**GG**A**ACT**G**CT**A**
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 T**A**CC**T**AC**G**CT**A**CC**T**C**G**TT**C**C**T**G**C**AG**A**AG**T**AC**A**T**C**AG**T**G**A**C**A**GG**A**TT
 G**A**AA**A**AT**G**A**A**GG**A**AT**A**AA**A**AG**A**AG**C**AA**A**AG**A**AT**G**T**G**A**A**AG**A**GC**A**AG**A**
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RG2C deduced polypeptide sequence (SEQ ID NO:92)

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RG2D polynucleotide sequence (SEQ ID NO:93) and (SEQ ID NO:94)

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AATACA (SEQ ID NO:93)

Sequence gap

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10 ACCATAGCAACTTGATAGGATGATTATTAAGAGAGAGTAAATATTATA
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 30 CTTAACCTAAATTATGTTGGTAAATGTTGGACAAGTATGGAAAATTA
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RG2D deduced polypeptide sequence (SEQ ID NO:95)

MAMETANEIIKQVVPVLMPINDYLRYVVSCRKYISDMMDLKMELKEAKDNVEE
 35 HKNHNISNRLEVPAQVQSWLEDVEKINAVENTVKDVGCCFNLKIRYRAGRDAF
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 LEANHMIALCGMGRVGKTHMMQRLLKVAKEKRKFQGYIIIVIGEISDPIAIQVVA
 DYLCIELKESDKKTRAEKLQRQFKAKSDGGNTKFLIILDDVVQSVLEDIGLSPSPN
 QGVDFKVLLTSRDEHVCVMGVEANSIINVGLLIEAEAQRLFQQFVETSEPELHKIG
 40 EDIVRRCCGLPIAIKTMACTLRNKRKDAWKDALSRQLQHHDIGNVATAVFRTSYENL
 PDKETKSFLMCGLFPEDFNIPTTEELMRYGWGLKLFDRTYTIIEARNRLNTCIERLV
 QANLLIGSDNGVHVKMHDLVRAFVLGMYSEVEQASIVNHGNMPGPWPDENDMIVH

SCKRISLTCKGMIEIPVDLKFPKLTILKLMHGDKSLKFPQEYEGMEKLQVISYDKM
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 5 DSKLNGLFLKTEVLFSLVHGMNDLEDVEVKSTHPTQSSFCNLKVRIISKCVELRYL
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 CHNVNIIGLPHLVDLKLKGIPGFTVIYPQNKLRTSSLKEEVVIPKLETLQIDGMENL
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 10 GCKRFRNIFTPITANFDLVALLEIHIGNYRENHESEEQIEILSEKETLQEVTDTNISND
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 LQELYLRNMDNTSHVWKCSWNKFFTLPKQQSESPFHNLTTIEMRWCHGFRYLFS
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 15 DALSSVIPCYAAGQMQLQVLTVSSCNGLKEVFETQLGTSSNKNNEKGCEEGIPR
 VNNNVIMPLNLKILEIYGCGLHEHIFTSALESRQLQELTIKYTLVNLPNLKEM
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 NCSLMEEVIVKDADVSVEEDKEKESDGKTNKEILVPLHLKSLKLQLRLSLKGFLGK
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 20 DFKQDSD.CEVNIK

RG2E polynucleotide sequence (SEQ ID NO:96)

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 25 TCAACAAGCTGTAGCGGATTACCTTGTATAGAGTAAAAGAAAGCACTAAACC
 AGCAAGAGCTGATAAGCTTCGTGAATGGTTAACGCCACTCTGGAGAAGGTA
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 40 ACCATGGTAATATGCTTGGATGGCCTGAAAATTATATGACCAACTCTTGC
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RG2E deduced polypeptide sequence (SEQ ID NO:97)

10 WEDTMMQRLKKVAKENRMFNYMVEAVIGEKTDPLAIQQAVADYLCIELKESTKP
 ARADKLRWFKANSSEGKNKFLVIFDDVWQSVDLEDIGLSHFPNQGVDFKVLLTS
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 15 VHVKMHDLVRAFVLVMFSEVEHASIINHGNMLGPENYMTNSCKTISLTCKSMSE
 FPGDLKFPNLTILKLMHGDKLLRYPQDFYEGMEKLWVISYDEMKYPLLPSLPQCSI
 NLRVLHLHRCSDLMMFDCCSICGNMLNLEVLSFKSGIEWLPSTIGNLKKRLLLDLRD
 CYGLRIEKGVKVLNVKIGGIYIGRADIL.

20 RG2F polynucleotide sequence (SEQ ID NO:98)

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 ATGTTAACCTTATTGTTGAAGCAGTTAGGGAAAAGACAGACAGCCCCGTTGCC
 ATTCAAGGATGCTATAGCAGATTACCTAGGTGTAGAGCTCAATGAAAATCTAAG
 CAAGCAAGAGCTGATAAGCTCCGTCAAGGATTCAAGGACAAATCAGATGGAGG
 25 CAA.AAATAAGTTCTTGTAAACTTGACGATGTTGGCAGTCTGTTGATCTGGA
 AGAATTGGTTAACGTCTTCCAATCAAGCGTCGACTTCAAGGTCTGTT
 GACATCACCGAGACAGACATGTTGCACAGTGATGGGGGTGAAGCAAATTAA
 TTCTAAACGTGGACTTCTAATTGAAGCTGAAGCACAAAGTTGTTCCACCAAT
 TTGTTGTCACCTCTGAGCCCGAGCTCCATAAGATAGGGAGAAGATTGTAAAGA
 30 AGTGTTCGGTCTGCCAATTGCCATCAAAACCATTGGCATGTACTCTACGACATA
 AAAGAAAGGATGCATGGAAGGATGCACTTCACGTTAGAGCACCATTGACATT
 CAAAGTGTGTGCCTAAAGTATTGAAACGAGCTACAACAATCTCAAAGACAA
 GGAGACTAAATCCGTATTTGATGTGTGGTTGTTCTGAAGACTTGGATAT
 ACCTATCGAGGAGTTGATGAGGTATGGATGGGGCTTAAGATTATTGATAGAGT
 35 TAATACTATTACACAAGCAAGAACACAGGCTCAACACCTGCATTGAGCGACTGG
 TGCACACAAATTGTTAATTGAAAGTGTGATGGTGTGCATGTCAAGATGCATG
 ATCTGGTTCGTGTCTTGTGTTGGAAATGTTTCTGAAGTGGAGCATGCTCAAT
 TGTCAACCATTGTAATATGCCGAGTGGACTGAAATGATATGACTGACTCTG
 CAAACAAATTCTTACATGCAAGAGTATGTTGGAGGTTCTGGAGACCTCAA
 40 GTTTCCAAACCTAAAGATTGAAACTTATGCATGGAGGTAAGTCACTAAGGTA
 TCCTCAAGACTTTATCAAGGAATGGAAAAGCTGGAGGTTATATCATAACGATGA
 AATGAAGTATCCATTGCTCCCTCGTTGCCTCAATGTTCCACCACCTCGAGTG

5 CTTCATCTCCATGAATGTTCATTAAGGATGTTGATTGCTCTCAATCGGTAAATC.
 TTTCAACATGGAAGTGCTCAGCTTGCTAATTCTAGCATTGAATTGTTACCTTC
 CGTAATTGAAATTGAAGAAGTGGCGCTGCTAGATTGACAAACTGTTATGG
 TGTCGTATAGAAAAGGATGTCCTGAAAAATTGGTGAAGAACTTGAAGAGCTTA
 TATTAGGAATGGTCTACCAGTTACAGAGGAT

RG2F deduced polypeptide sequence (SEQ ID NO:99)

10 VEDTMMQLKKVVHEKKMFNFIVEAVIGEKDPVAIQDAIADYLGVELNEKSQQA
 RADKLRQGFKDSDGGKNKFFVILDDWQSVDLEDIGLSPFPNQGVDFKVLLTSRD
 RHVCTVMGVEAKLILNVGLLIEAEAQLSFHQFVVTSEPELHKIGEDIVKKCFGLPIAI
 KTMACTLRHKRKDAWKDALSLEHHDIQSVPVKVFETSYNNLKDKEKSVFLMCG
 LPEDLDIPIEELMRYGWGLRLFDRVNTITQARNRLNTCIERLVHTNLLIESVDGVH
 VKMHDLVRASFVLGMFSEVEHASIVNHGNMPEWTENDMTSCKQISLTCKSMLEFP
 15 GDLKFPNLKILKLMHGGKSLRYPQDFYQGMEKLEVYSISDEMKYPLLPSLPQCSTILR
 VLHLHECSLRMFDCSSIGNLFNMEVLSFANSIELLPSVIGNLKKLRLLDLNCYGV
 RIEKDVLKNLVKLEELYIRNGLPVYRG

RG2G polynucleotide sequence (SEQ ID NO:100)

20 GAAGACACGATGATGAAGAACTGAAGGAGGTCGTGGGACAAAGAAATCATTCA
 AATATTATTATTCAAGTGGTCATAGGAGAGAACAAACCTATTGCAATTCTAG
 CAAGCTGTAGCAGATTACCTCTATAGAGCTGAAAGAAAACACTAAAGAAC
 AAGAGCTGATAAGCTTCGAAACGGTTGAAGCCGATGGAGGAAAGAATAAGT
 TCCTTGTAAACTTGACGATGTATGGCAGTTGTCGATCTGAAGATATTGGTT
 25 AAGTCCTCTGCCAAATAAAGGTGTCAACTTCAAGGTCTTGTGACGTCAAGAGA
 TTACATGTTGCACTCTGATGGGAGCTGAAGCAAATTCAATTCTTAATATAAA
 AGTTTAAAAGATGTAGAAGGACAAGTTGTCGCCAGTTGCTAAAATGC
 GGGTGATGATGACCTGGATCCTGCTTCAATGGATAGCAGATAGTATTGCAAG
 TAGATGTCAAGGTTGCCATTGCCATCAAACCATGCCCTAACGTTAAAGG
 TAGAAGCAAGTCTGCATGGGACGTTGCACTTCTGCTGGAGAACATCAAAGAT
 30 TGGTAGTGAAGAAGTTGTCGTGAAGTTTAAAATTAGCTACGACAATCTCCA
 AGATGAGGTTACTAAATCTATTTTACTTTGTCCTTATTCTGAAGATT
 GATATTCCACTGAGGAGTTGGTGAGGTATGGTGGGCTGAAATTATTATA
 GAAGCAAAACTATAAGAGAACAGGCTCAACACCTGCACTGAGCG
 GCTTAGGGAGACAAATTGTTATTGGAAGTGTGACATTGGATGTGTCAGAT
 35 GCACGATGTGGTGCCTGATTTGTCATATATTCTCAGAAGTCCAACACGC
 TTCATTGTCAACCATTGTAACGTGTCAGAGTGGCTAGAGGAAAATCATAGCAT
 CTACTCTGTAAAAGAATTCTTACATGCAAGGGTATGTCTCAGTTCCCAA
 AGACCTCAAATTCCAACCTTCAATTGAAACTTATGCATGGAGATAAGTC
 ACTGAGCTTCCCTGAAAACCTTATGGAAAGATGGAAAAGGTTCAGGTAATATC
 40 ATATGATAAAATTGATGTATCCATTGCTCCCTCATCACTTGAATGCTCCACCAA
 CGTTGAGTGCTTCATCTCATTACTGTCATTAAGGATGTTGATTGCTCTCA
 ATTGGTAATCTCTCAACATGGAAGTGCTCAGCTTGTAAATTCTAACATTGAA

TGGTTACCATCTACAATTGGAAATTGAAGAAGCTAAGGCTACTAGATTTGACA
 AATTGTAAAGGTCTCGTATAGATAATGGTGTCTAAAAAATTGGTCAAACCTT
 GAAGAGCTTATATGGGTGTTAACGTCCGTATGGACAGGCCGTAGCTTGACA
 GATGAAAA

5

RG2G deduced polypeptide sequence (SEQ ID NO:101)

RHDDEELKEVVGQKKSFNIIQVVIGEKTNPIAQAVADYLSIELKENTKEARADKL
 RKRFEADGGKNKFLVILDDVWQFVDLEDIGLSPLPNKGVNFKVLLTSRDSHVCTL
 MGAEANSILNIKVLKDVEGQSLFRQFAKNAGDDDDLDPAFNGIADSIAASRCQGLPIAI
 10 KTIALSLKGRSKSAWDVALSRLENHKIGSEEVVREVFKISYDNLQDEVTKSIFLLCAL
 FPEDFDIPTIELVRYGWGLKFIEAKTIAREARNRLNTTERLRETNLLFGSDDIGCVK
 MHDVVRDFVLHIFSEVQHASIVNHGNVSEWLEENHSIYSCKRISLTCKGMSQFPKDL
 KFPNLSILKLMHGDKSLSPENFYGKMEKVQVISYDKLMPPLLSSLECSTNVRVLH
 15 LHYCSLRMFDCCSIGNLLNMEVLSFANSNIEWLPSTIGNLKKLRLLDLTNCGLRID
 NGVLKNLVKLEELYMGVNRPYGQAVSLTDE

RG2H polynucleotide sequence (SEQ ID NO:102)

TGAAGGAGGTTGTGGAACGAAAGAAAATGTCAGTATTATTGTTCAAGTG
 GTCATAGGAGAGAACAAACCTATTGCTATTCAAGCAAGCTGTAGCAGA
 20 TTACCTCTATAGAGCTGAAAGAAAACACTAAAGAACAGCAAGAGCTGATA
 AGCTTCGTAATGGTCGAGGCCGATGGAGGAAAGAACAGTCTCTTGTA
 ATACTTGACGATGTATGGCAGTTGTCATCTGAAGATATTGGTTAAG
 TCCTCTGCCAATAAAGGTGTCACCTCAAGGTCTGTGACGTCAAGAG
 ATTACACATGTTGCACTCTGATGGGAGCCGAAGCCAATTCAATTCTCAAT
 25 ATA.AAAGTTAACAGCTGTAGAAGGACAAAGTTGTTCCGCCAGTTGC
 TAAAATGCGGGTGATGATGACCTGGATCCTGCTTCAATAGGATAGCAG
 ATAGTATTGCAAGTAGATGTCAAGGTTGCCATTGCCATCAAACCAATT
 GCCTTAAGCTTAAAGGTAGAACAGCAAGCCTGCGTGGGACCATGCGCTTTC
 TCGTTGGAGAACCATAGATTGGTAGTGAAGAAGTTGTGCGTGAAGTT
 30 TTA.AAATTAGCTATGACAATCTCAAGATGAGATTACTAAATCTATTTT
 TTACTTGTGCTTATTCCTGAAGATTGATATTCTACTGAGGAGTT
 GATGAGGTATGGATGGGCTTGAAATTATTAGAACAGCAAAACTATAA
 GAGAACAGCAAGAACAGGCTAACACACTGCACTGAGCGGCTAGGGAGACA
 AATTGTTATTGGAAGCGATGACATTGGATGCGTCAAGATGCACGATGT
 35 GGTGCGTGATTTGTTGCATATATTCTCAGAACAGTCCAGCACGCTCAA
 TTGTCAACCATTGTAACGTGTCAGAGTGGCTAGAGGAAATCATAGCATC
 TACTCTGTAAAAGAATTCAATTAAACATGCAAGGGTATGTCTGAGTTCC
 CAAAGACCTCAAATTCCAAACCTTCAATTGAAACTTATGCATGGAG
 ATAAGTCGCTGAGCTTCCGTAAAACATTGAAAGATGGAAAAGGTT
 40 CAGGTAATATCATATGATAAATTGATGTATCCATTGCTCCCTCATCACT
 TGAATGCTCCACTAACGTTCGAGTGCCTCATCTCCATTATTGTCATTAA
 GGATGTTGATTGCTCTCAATTGGTAATCTCTCAACATGGAAGTGCTC

AGCTTGCTAATTCTAACATTGAATGGTACCATCTACAATTGGAAATT
 GAAGAAGCTAAGGCTACTAGATTTGACAAATTGTAAAGGTCTCGTATA
 ATAATGGTGTCTAAAAAATTGGTCAAACTTGAAGAGCTTATATGGGT
 GTTAATCATCCGTATGGAC

5

RG2H deduced polypeptide sequence (SEQ ID NO:103)

KEVVERKKMFSIVQVVIGEKTNPIAIQQAVADYLSIELKENTKEARADKLKWFEA
 DGGKNKFLVILDDVWQFDLEDIGLSPLPNKGVNFKVLLTSRDSHVCTLMGAEAN
 SILNIKVLTAVEGQSLFRQFAKNAGDDDLDPAFNRIADSIASRCQGLPIAIKTIALSLK
 10 GRSKPAWDHALSRLENHKIGSEEVVREVFKISYDNLQDEITKSIFLLCALFPEDFDIP
 TEELMRYGWGLKLFIEAKTIAREARNRLNTCTERLRETNLLFGSDDIGCVKMHDVVR
 DFVLHIFSEVQHASIVNHGNVSEWLEENHSIYSCKRISLTCKGMSEFPKDLKFPNLSI
 LKLMHGDKSLSPENFYGKMEKVQVISYDKLMYPLPSSLECSTNVRVLHLHYCSL
 15 RMFDCSSIGNLLNMEVLSFANSNIEWLPSTIGNLKKRLLDLTNCKGLRIDNGVLKN
 LVKLEELYMGVNHPYG

RG2I polynucleotide sequence (SEQ ID NO:104)

AAGAAGAGCTGAAGGAGGTTGTGGAACAAAAGAAAACGTTCAATATTATT
 GTTCAAGTGGTCATAGGAGAGAAGACAAACCTATTGCTATTCAAGCAAGC
 20 TGTAGCAGATTCCCTCTCTATAGAGCTGAAAGAAAACACTAAAGAAGCAA
 GAGCTGATAAGCTTCGAAATGGTCGAGGCTGATGGAGGAAAGAATAAG
 TTCCCTCGTNATACTTGACGATGTATGCCNGTTGATCTGAAGATAT
 TGGTTAACGCCTCATCCAAATAAAGGTGTCANCTCAAGGTCTTGTGA
 CGTCAAGAGATTCACATGTTGCACTCTGATGGGAGCTGAAGCCAATTCA
 25 ATTCTCAATATAAAAGTTAAAAGATGTAGAAGGAAAAGTTGTTCCG
 CCAGTTGCTAAAATGCGGGTGTGATGACCTGGATCTGCTTTCATTG
 GGATAGCAGATAGTATTGCAAGTAGATGTCAAGGTTGCCATTGCCATC
 AAAACCATTGCCTTAAGTCTAAAGGTAGAAGCAAGTCTGCATGGACGT
 30 TGCACTTCTCGTCTGGAGAACATCAAAGATTGGTAGTGAAGAAGTTGTGC
 GTGAAGTTTAAAATTAGCTATGACAATCTCAAGATGAGGTTACTAAA
 TCTATTTTTACTTTGTGCTTTATTCCTGAAGATTGATATTCTAC
 TGAGGAGTTGGTGTGAGGTATGGTGGGGCTTGAAATTATTAGAAGCAA
 AAACTATAAGAGAACAGAAACAGGCTCAACACCTGCACTGAGCGGCTT
 35 AGGGAGACAAATTGTTATTGGAAGTGTGACATTGGATGCGTCAAGAT
 GCACGATGTGGTGCCTGATTGTTGCTATATTCTCAGAAGTCCAGC
 ACGCTTCAATTGTCAACCATGGTAATGTGTCAAGAGTGGCTAGAGGAAAAT
 CATAGCATCTACTCTGTAAAAGAATTCTATTAAACATGCAAGGGTATGTC
 TGAGTTCCAAAGACCTCAAATTCCAACCTTCAATTGAAACTTA
 40 TGCATGGAGATAAGTCGCTGAGCTTCCTGAAAACCTTATGGAAAGATG
 GAAAAGGTTCAGGTAAATCATATGATAAATTGATGTATCCATTGCTTCC
 CTCATCACTGAATGCTCCACCAACCTTCGAGTGCTCATCTCCATGAAT
 GTTCATTAAGGATGTTGATTGCTCTCAATTGTAATCTCTAACATG

GAAGTGCTCAGCTTGCTAATTCTGGCATTGAATGGTTACCATCTACAAT
TGGAAATTGAAGAAGCTAACGGCTACTGGATCTGACAGATTGTGGAGGTC
TTCATATAGATAATGGCGTCTAAAAAATTGGTCAAACCTGAAGAGCTT
TATATGGGTGCTAACCGTCTGTTGGAAAGTGCCAT

5

RG2I deduced polypeptide sequence (SEQ ID NO:105)

EELKEVVEQKKTFNIVQVVIGEKTNPIAIQQAVADSLSIELKENTKEARADKLKWF
EADGGKNKFLVILDDVW?FVDLEDIGLSPHPNKGV?FKVLLTSRDSHVCTLMGAEA
NSILNIKVLDVEGKSLFRQFAKNAGDDLDPAFIGIADSIAsrcQGLPIAIKTIALSL
10 KGRSKSAWDVALSRLENHKIGSEEVVREVFKISYDNLQDEVTKSIFLLCALFPEDFDI
PTEELVRYGWGLKLFIEAKTIAREARNRLNTCTERLRETNLLFGSDDIGCVKMHDVV
RDFVLHIFSEVQHASIVNHGNVSEWLEENHSIYSCKRISLTCKGMSEFPKDLKFPNLS
15 ILKLMHGDKSLSPENFYGKMEKVQVISYDKLMYPLLSSLECSTNLRLLLDCGGLHIDNGVLKN
LVKLEELYMGANRLFGKCH

RG2J polynucleotide sequence (SEQ ID NO:106) and (SEQ ID NO:107)

ATGTCCGACCCAACAGGGATTGTTGGTGCATTATTAAACCAATTGCTCA
AACGGCCTGGTCCCCTACAGACCATGTAGGCTACATGATTCCCTGCA
20 GAAAATATGTGAGGGACATGCAAATGAAAATGACAGAGTTAAATACCTCA
AGAACATCAGTCAGAGGAACACATTAGCCGAACACAAGAAATCATCTTCA
GATTCCATCTCAAATTAAAGGATTGGTGGACCAAGTAGAGAAGGGATCAGAG
CGAATGTTGCAAACCTTCCAATTGATGTCATCAGTTGTTAGTCTCAGG
ATCAGGCACAAGCTTGGACAGAAAGCCTCAAGATAACTGAGCAGATCGA
25 AAGTCTAACGAGACAAAATCGCTGATTATCTGGACTGATGAACCTGTT
CCCTGGGAAGAGTTGGTCCATGATTGCATCCACCTCTGCAGCATCAAGT
GATCATCATGATGTCTCCCTCAAGAGAGCAAATTTTAGGAAAGCACT
AGAACGACTTGAACCCGTCAAAACCCACATAATAGCCTATGGGGGA
TGGCGGAGTGGGAAGACCACGATGATGAAGAAGCTGAAAGAGGTCGTG
30 GAACAAAAGAAAACGTGCAATATTATTGTTCAAGTGGCATAGGAGAGAA
GACAAACCTATTGCTATCCAGCAAGCTGTAGCAGATTACCTCTATAG
AGCTGAAAGAAAACACTAAAGAAGCAAGAGCTGATAAGCTTGTAAACCGG
TTCGAAGCCGATGGAGGAAAGAATAAGTCCCTGTAATACTGACGATGT
ATGGCAGTTTCGATCTTGAAGATATTGGTTAAGTCCTCTGCCAATA
35 AAGGTGTCAACTCAAGGTCTGTTGACGTCAAGAGATTCACATGTTGC
ACTCTGATGGAGCTGAAGCCAATTCTATTCTCAATATAAAAGTTTAAA
AGATGTAGAAGGAAAAGTTGTTCCGCCAGTTGCTAAAATGCGGGTG
ATGATGACCTGGATCCTGCTTCATTGGGATAGCAGATAGTATTGCAAGT
AGATGTCAAGGTTGCCATTGCCATAAAACCATTCGCTTAAGTCTAA
40 AGGTAGAAGCAAGTCTGCATGGGACGTGCACTTCTCGTCTGGAGAATC
ATAAGATTGGTAGTGAAGAAGTTGCGTGAAGTTTAAAATTAGCTAT
GACAATCTCCAAGATGAGGTTACTAAATCTATTTTACTCTGTGCTTT

ATTCCTGAAGATTGATATTCTATTGAGGAGTTGGTGGGT
GGGGCTGAAATTATTATAGAAGCAAAACTATAAGAGAAGCAAGAAC
AGGCTAACAACTGCACTGAGCGGCTAGGGAGACAAATTGTTATTGG
AAGTCATGACTTGGGTGCGTCAAGATGCACGATGTGGTGCCTGATTG
5 TTTGCATATGTTTCAGAAGTCAGCATGCTCAATTGTCACCAGGT
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ATCCAACACCTTGATTGAAACTTATGCATGGAGATAAGTCGCTGTGC
TTCTGAAAACCTTGATTGAAAGATGGAAAAGGTTAGGTAATATCATA
10 TGATAAATTGATGTATCCATTGCTTCCCTCATCACTGAAATGCTCCACTA
ACGTCGAGTGCTTCATCTCATTATTGTCATTAAGGATGTTGATTGC
TCTTCATTGTAATCTCTCACATGGAAGTGCTCAGCTTGCTAATT
TAACATTGAATGGTTACCATCTACAATTGAAATTGAAAGAAGCTAACGC
TACTAGATTGACAAATTGTAAGGCTTCGTATAGATAATGGTGTCTTA
15 AAAAATTGGTCAAACCTGAAGAGCTTATATGGGTGTTAATCGTCCGTA
TGGACAGGCCGTTAGCTGACAGATGAAAATGCAATGAAATGGTAGAAG
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GGGATGTTCTTACATGGATCTTCAGTAAAAGCAGGCACTCATACGAAA
20 ACACGTTGAAGTGGCATTGACAAAGGCGAATATTGGAATCCGAATG
AACGGGTTGTTGAGAAAACGGAGGTTCTTGTAAAGTGTGGGGATAT
GTATCATCTTCAGATGTTAAGGTGAAGTCCTCTCGTTACAATTAA
GAGTCCTGTCGTTAGAGTGTGCAGAGTTGAAACACCTCTCACACTT
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25 CGATAATATGGAAGAACTCATACATACCGGGGTAGTGAAGGAGATA
TTACATTCCCCAAGCTGAAGCTTATATTGATGGCTGCCAACCTA
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30 TATGTTAACACATTAAACAATCTTCAACTAAAGTTCAGAACATATA
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ATGACATGGAGAATTAAAGGAAATATGGCCTAGTGAGCTTAGAGGT
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35 GAATCTATTCCACACAATCCCAGTCTCTGCTGCATCATCTGAAGAGC
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CATCAATGTGGAGAATTCAATGAAGCTAACAGAGGAGGTGAGGATAAAAG
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40 ATAATCATTACGAGATGTAAGAGGTTACAAATGTATTCACACACCTATCAC
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10 TNGTNTCACAAAGGGATATATAGTAAAATATTATTTGCAGTCAT
GCATAGTTGTATTTAAATGATTATAACGTTGAGTGGAAACCA
CTCAATCTAGTAGACCCACTATCACATGTACATCAGCTTACATCTATT
TTCTTCTCCTTTTCATCTTTAAACTCATAACACNTAAAANTANC
ATATTTCCAACACACTNAACTCATTGTACACATTATTATTTAATTAA
15 TTA-AATTNGAAAATTAAAATTAANTAAANCNTAACATTTTAATTAAAA
AATATTAATCCAATAAAANTNCACGATAAATTAAAANGTTANTTG
GAAAANCC (SEQ ID NO:106)

Sequence gap

ATAACCCTTCAGGGTCAACTCAAGTCCAAGTTAAAGTCAGGTCAAAA
20 CCTTGGTAAAGTCACCTGGTCAAAGTCACATCTACTGACTCACCT
CACCGAGTTGGTCCACCAACTGTGCGAGTCCCTTAATCCACAAACTTCAA
GAACCTCGATCCTACTCGTCAGTCTTCAGAAACTCTCGAGTTCCAT
TACACAGAATCGGGACCTTTGCTCATGACTCGCCGAGTTCATCCTTGAA
CTTGTGAGTCTAGCTCATACGAGTCAGTGTAGTCCTGACTCGT
25 CGAGTTCTCCTTGAACTCGTCAGTCCATCTCGTATAGTTGGGACATT
GCCTTGAACTCACCGAGTTCATCATTGAACTCATCGAGTCCTCGATCTT
CAAGTCCATAATCCTGTCATCTGTGAGTCCTCTAGACTCAACCA
GATTCCCTAGAAACAGAAAAGGTTAGGGAACCAATTACCTGACTGCCGAG
TCCCAAGAACGAATCCCCGAGTCCCCAATGTCCATGACCATAATCGA
30 TTTCTGTTGGCTCATTGCATCCAAAGCATAGATCTAACCTCTAGGGTC
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CATAAAGTTAACACCTTATGCCATGGGAATCTCAATGGTCCATATCT
GAAGTTAACACTCTACAATATGTTCAAACCGAAGGTGGCTAGAAATG
35 CCCCAAAATGGCAAGATTCAAGCCTAAAGGAGATCTAACAAATGATAAG
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25 CCTCCAGCATTGGATCTAACGGGTATGGACAACATGATTGCGTGTGGA
AGTGCAGCACTGGAATAAATTCTTCACTCTTCCAAACAACATCAGAA
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TAAGTACTGTTTCACCTCTATGGCAGAACTTCTTCCAAACCTAAAGA
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30 GATGATGAGGATGAAGAAATGACTACATTACATCTACCCACACAACCAC
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5 AATACGCTAGAGAGATAAGTATAGAATTCTGCAATGCATTGTCAAGTGTG
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10 GGAAATCAGCTTGTGGGGTTGGAACATATATTCACATTCTGCAC
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CTTCT
15 TTTCTTGTCTAAAGTCATTGTATTGGTCAATCTACCAGAGCTGGTAGG
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GCTCCCCAACTCAAGTATATACACACAAGATTAGGCAAACATACTATTGA
TCAAGAATCTGCCCTAACCTTCATCAGGTATATATGTTCTTAATTGG
20 CATCATCTAATTAAAGAAAGATATCATTCTGCCAAGTAAATTACTCAA
ACACATTCACACTGGTTTCAGTCTAAGTTATGTTCTAGGAAGGCCA
AAATGGGAAAGCAAGATAGGGAAAAATAGTGTATTGAGCTGGAAAGGGTA
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25 CATTCTTAGCCTCTCGAACAGCTAGAAACCCTTAATCTTTGAT
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 T (SEQ ID NO:107)

25

RG2J deduced polypeptide sequence (SEQ ID NO:108)

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 15 SIIKIKQQDFKKAQDSI.CEVNTR

RG2K polynucleotide sequence (SEQ ID NO:109) and (SEQ ID NO:110)

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35 AAACAGATGTTCTTATTAAAGTGTGGGAGATATGAATGATCTTGAAGAT
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40 AACCTTGAGTTATTGGATATAAGTTTATGGACAGCATGAGTCATGTATG
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AAGGTATGTTTTTTNTNCCCTT (SEQ ID NO:109)

Sequence gap

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 5 GATTGCCACAAATGATGGGTTCACACCTGGTGGTCAACAACTTCCA
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 (SEQ ID NO:110)

10 **RG2K deduced polypeptide sequence (SEQ ID NO:111)**

MECITGIFSNPFAQCLIAPVKEHLCLLIFYTQYVGDMLTAMTELNAAKDIVEERK
 NQNVEKCFCVPNVNRWLEDVQTINRKVERVLNDNCNFNLCNRYMLAVKAL
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 15 ISIQDAVADYLDMKLTESNESERADKLREGFQAKSDGGKNRFLIILDDVVQSVN
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 IETIAHVVFQMSYDNLQNEEAQSIFLLCGLFPEDFDIPEELVRYGWGLRVFNGV
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 20 NHNGGGMLGWPENDMSASSCKRISLICKGMSDFPRDVFPNLLILKLMHADKS
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 25 VEVKLAHLPKSSSFHNLRLVIISECIELRYLFTLDVANTLSKLEHLQVYECNMEEII
 HTEGRGEVTITFPKLKFLSLCGLPNLLGLCGNVHIINLPQLTELKNGIPGFTSIYPEK
 DVETSSLLNKEVVIPNLEKLDISYMKDLKEIWPCELGMSQEVDVSTLRVIKVSSCDN
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 30 LSEVWRIKGADNSSLLISGFQGVESIIVNKCKMFRNVFTPTTNFDLGALMEIRIQDC
 GEKRRNNELVESSQESEQ

RG2L polynucleotide sequence (SEQ ID NO:112)

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 5 GAGGAGTTGATGAGGTATGGATGGGCTTAAAGCTATTGACAGAGTTA
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 20

RG2L deduced polypeptide sequence (SEQ ID NO:113)

EDTMMQRLKKVAKENRMFSYMVEAVIGEKDPIAIQQAVADYLRIQFESTKPAR
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 HVCTMMGVEANSVINVGLLTEVEAQSLFQQFVETFEPELCKIGEVIVRKCCGLPIAI
 25 KTMACTLRNKRKDAWKDALSRIEHYDIRSVAPKVFETSYHNLQDRETKSVFLMCG
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 30 YGLCIEQGVNLVELEELYIGNASAFRDYN岑NEMA

RG2M polynucleotide sequence (SEQ ID NO:114)

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 AGTTTGTGGAGACCTNAANTTCCAACCTAATGATTAAAACCTATG
 CATGGAGATAAACGCTAAGGTT

RG2M deduced polypeptide sequence (SEQ ID NO:115)

15 GEDTIDAKAEEVAKEKRMFSYIIIEAVIGEKTDPISQEAIISYLGVELNANTKSVRAD
 MLRQGFKAQSDVGDKFLIILDDVWQSVLEDIGLSPFPNQGVNFKVLLTSRDRHI
 CTVMGVEGHSIFNVGLLTEAESKRLFQFWQFVEGSDPELHKIGEDIVSKCCGLPIAKT
 MACTLRDKSTDWKDALSRLEHHDIENVASKVFRASYDHLQDEETKSTFFLCGLFP
 20 EDSNIPMEELVRYGWGLKLFKKVYTIREARTRLNTCIERLIYNLLIKVDDVQCIKM
 HDLIRSFVLDMFSKVEHASIVNHGNTLEWPAD??HDSCKGLSLTCKG?CEFCGDL?F
 PTLMILKLMHGDKSLRF

RG2N polynucleotide sequence (SEQ ID NO:116)

25 AGGTAAAATCCATAACCCTAAATGTTGGTACGCTCATATATCAAATTGCG
 TGTGTTGTTGAATGAAAAAAAGCATGCTCAAAAACCAGTGTAAAGGCACGG
 TATATGACATATTATAGTTACTGATAACAAATTATGATAATTGGGTT
 TACRTAAGTTAGGATTCGTACTTCACCAAATGTAATAGTTTGTGAGT
 CTATCTATGTATTGGGAATCACATTAGCAACGGGATTGTACTAGTAAT
 TCGAAAAAGTCTTTAAATAATTCTGTTATAATTATGAATAGTT
 30 TAGCGACATCTAATATTAAAGAATGTATCTGATATTGAATTAATGTCC
 TTAATGTGAACATAGACCTTCCATTACTAATGCCTAATTATTAGTT
 CTAATCAATAAATTAAATTCTGTTATGCTCTAAGACAATAAAAT
 CCATGATTACCTTAAATATTAACAAAAATGACCATAAAATAAAAT
 ATTAGGATACCAACCCCCCGCCATGCCAATGTCTAAATATTCTGAT
 35 GCTTTGCTTCCCTTTCTGTTAGTCTATTATTCTGGAGAGTT
 GAGAGAGTTCATACAAGAAAATTCAAGAAGAAAGCAAAGGTCCAGGT
 TTCTCTTCTTAATTATGTATTAACCTACAAGCATTACAGATCC
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 CCTTGATGATTCATAGATATTGCATGGAGTTGAGATTGTAAAGAAAAG
 40 TGGTGAATAGAAAGAGCAAGTGAATCCAGATATAGTATTGGTAATATG
 ATGATGAGATAGAGATATGTTAAAACGGCTAGAAAATTGTTAATTG
 AAATTAGGTGTTGAATTGAAAGATAACCAAGCTAATAACTAATTAGTT

ATGCTAAWTAGTTATAAAGAACAAACACTCTAGTTTTTTTTCATGA
TTTCAACCTCTTGACCAAACAAATTATAGCAAAATTGAATATCATT
CTCTGCAATCAATCTAACCTTTGTTATTATCATCATGTCTAAAATTGCC
ACAAGTTATTTCAAAGTCATATTGGATTATGAAAGGACTATTTTACC
5 AATTACATCTTACTTTATGGGCCAAGCTAATACAATCCGACTAAACTA
AAGGAATATGGGATGCATATAGTTGCTTCCCATTAGATTTCTATCT
AATTTGTCTATTGTAATTAGGTGCCACCAAGTAAATTGTTAAA
TGGATATCGTTAATGCCATTCTAACCAAGITGTCGAGACTCTCATGGTA
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10 GGAAATGGGTATCAAATGAGGGATTGAATGCTACTAGACTTGGTGTGCG
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TTTCCTAGCGATGTTGGCAGTTGTTCAATCTAACGTTAGACACGGGG
TCGGAAAGAGAGCCTCCAAGATAATTGAGGACATCGACAGTGTATGAGA
15 GAACACTCTATCATCTGGAATGATCATTCCATTCTCTAGGAAGAAT
TGATTCCACGAAAGCATCCACCTCAATACCATCACCGATCATGATG
AGTTCCAGTCAGAGAGCAAACCTTCACAGAACGACTAAACGCACTCGAT
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GAAGACGACAATGATGCATCGGCTGAAAAGGTTGTGAAAGAAAAGAAAA
20 TGTTTAATTTATTGTTGAGGCGGTTGAGGGAAAAAACAGACCCCCATT
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CTGCTGGTAAGAAGATCCTAGTCATACTCGACGATGTATGGCAGTTGTA
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25 CAAGGTGTTGTTGACATCACGAGACAAAGATGTTGCACTGAGATGGGAG
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30 TGGAGAAATGCACTTCTCGTTAGTGAACTACAACATTGAAAATATAGT
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GTACTATAGGAGAACAGAACATCAGGCTAACACATGCATTGAGCGGC
35 TCATTACACAAATTGTTGATTGAAGTTGATGATGTTAGGTGCATCAAG
ATGCATGATCTGTCGTGCTTGTGTTGGATATGTATTCTAAAGTCGA
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40 GCATGAAGATATCATTGAGGTTCCAAAAACTTTATGAAGAAATGG
AGAAGCTTGAGGTTATATCCTATGATAAAATGAAATATCCATTGCTTCCC
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AAGTGCTTAGCTTGCTGATTCTGCCATTGACCTGTTGCCCTCCACAATC
5 GGAATTTGAAGAAGCTAAGGCTACTGGATTGACAAATTGTTATGGTCT
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10 ATAAAGAGCAGGCACTCGTATGAAAACACATTGAAGTTGGTTATTGACAA
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AAGTCATCCTCACAAACYTCTTCATCTCTCGTTCAACAATTAAAGAGT
CCTTGTGTTCAAAGTGTGCAGAGTTGAAACACACTTCTCACACCTGGTG
TTGCAAACACTTAAAAAAGCTTGAGCATCTGAAGTTACAAATGTGAT
AATATGGAAGAACTCATACGTAGCAGGGTAGTGAAGAAGAGACGATTAC
15 ATTCCCCAAGCTGAAGTTTATCTTGTGTTGCTACCAAAGCTATCGG
GTTTGTGCGATAATGTCAAAATAATTGAGCTACCACAACTCATGGAGTTG
GAACTTGACGACATTCCAGGTTACAAGCATATATCCATGAAAAAGTT
TGAAACATTAGTTGTTGAAGGAAGAGGTAATATAAATTAAATTGCT
AATACATTACAAAGGATCTTCAGTTAAATCTTCAAAATATATTGAA
20 TTTGATTGTATGGGTATTATTGTTGGATGGACTATTAAATAATGATTA
TCTTGCAGGTTCTGATTCTAACAGTTAGAGAAACTGCATGTTAGTAGTATG
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TAAGTTCAGAGAGATTAAGTGAAGTACTGTGATAAGCTGTGAATTGTT
TTCCGCACAAGCCCATACTCTGCTGCGTCATCTGAAGAGCTAAAGTC
25 AAGAATTGTGGTCCATTGAATCGTTATTCAACATCCATTGGATTGTGC
TGGTGCACACTGGAGATGAATACAACACAGTGGTGTAAAGAATTATTAAAG
TGATCAGTTGTGATAAGCTGTGAATCTCTTCCACACAATCCATGTCT
ATACTGCATCATCTGAAGAGCTTGAAGTCGAGAAATTGTTGCTGGATTG
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30 ACAACAGAACGCTAACAGAAACATCAAAGTGGAGAATTAGGAAAGCTA
AGAGAGGTGTGGAGGATAAAAGGTGGAGATAACTCTCGTCCCCCTGTTCA
TGGCTTCAATCTGTTGAAAGCATAAGGGTTACAAATGTAAGAGGTTA
GAAATGTATTCACACCTACCACCAAATTAAATCTGGGGGACTTTG
GAGATTCAATAGATGACTGCGGAGAAAACAGGGAAAATGACGAATCGGA
35 AGAGAGTAGCCATGAGCAAGAGCAGGTAAAGGATTCAATTCACTTCKT
ACTTAATTAATGATTAAGCTCTGTTTTRAATAAAAAGGGACAAACC
ATTTCATGACTTAATGTAGCAATACAAGTCATGTATAAGAGTGACCAACT
CTTTTTATTATAAAATGACTACAAATATTTCATTAGAGATCA
TGTATAATGTGACTAATTTCATCACCTAACCTTGTGATAATTCTT
40 TATAAATGTCACTAGTTACTTTCAGTAAAATAACAAATTAAATAAATTAA
TCAACAAAAAGCATCAACTAAAAAATCCCACAACCCGTAATAATTAAA
ATAAAAGGATTAAACATCTAACGAAACAATTCTAAACATGATT
TGGACCAAATACCAACAGCAACTCAAGTTGGAATCGATTGCTTAAAAA

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5 AAAATGGCTAGGTGGTCTGGTAGATGAAGATGGAAGGGAGA
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10 ACCAAAAGCGAACAAAAGTAAACAGTAGGGACCATCCGATTTAAAAAT
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15 TTTGTTATCTAGGTGTGACAAAAAAACGATAGTACCATGATGTGAACA
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GGAAACAGTTCCGTGAGACCGTGACTGGATGGTAGATAAATTAGTAAA
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20 CTTGTTGCATAGGATCCTAGCATCTTTAATAATTATTGAAGGTGAA
AGATCCAACATTAAATCTGTTGGCATTTCCATCATTGCAACTGTT
TCTTGAAAAAAA::TACCTAAAATCAAACCACTTCATATCCAAAA
TTATAAGAGAGAATTGTTAACGGACATGGAATCATAAATCATTAAACACAG
TTCAGTACACAGGTTGCTAATTACATTCTGCTGTGCAGATTGAAATT
TATCAGAGAAAGAGACATTACAAGAAGGCCACTGGCAGTATTCAAATATT
25 GTATTCCCATTCTGCTCATGCACTCTTCTATAACCTCCATAAACTTAA
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30 AAGAACAACTCAGAACCCCCATTCCACAAACCTCAGTAACACATACATATTAT
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35 :::GCACACAAACCAACCACTTTTCCCTCATCTGATTCTCACTCTAA
GCAACTGAAGAATCTGAAGTGTATTGGTGGAGGTGGCCAAGGATGAGG
GGAGCAATGAAATATCTTCAATAATACCACTGCAACTACTGCTGTTCTT
GATCAATTGAGGTATGCTTGTACATATTCAATTATTATTAATTCC
TTGTTAATTCTTCTTCTTGCATATTCTATGAAAAAAATCACCAA
40 TCACAAATAAGAGATTAAACTTTATTCAACCCATGCGGACTCAAGA
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TTTATTGTTATTATCATTCTCATATTACTGATAACATTCTTT
TTACTTTCTAATTAGAAAAGGTCCACATGTCTAATTAGGTTCCATTIC
TATGTGAATCCTCTATTCTGTCTGTAATCAAGCATCTAGATTATTATC

CATTTCTATAATTGTGTTATATTGACAGTTTTCTTTTATAGTTGT
 AATTGCAACCTGTCATATWTTMWWKKCWWWATKYWMWWARTAATAACATTT
 TATAACCCWCTATACTAAGATA

5 **RG2N deduced polypeptide sequence (SEQ ID NO:117)**

LGKTTMMHRLKKVKEKKMFNFIVEAVGEEKTDPIAQSAVADYLGIELNEKTKPA
 RTEKLRKWFDNSAGKKILVILDDVWQFVDLNDIGLSPLPNQGVDFKVLLTSRDKD
 VCTEMGAEVNSTFNVKMLIETEAQSLFHQFVEISDDVDRELHNIGVNIVRKCGGLPI
 VIKTMACTLRGKS KDAWNALLRVNYYNIENIVNGVFKMSYDNLQDEETKSTFLL
 10 CGMFPEDFNPTEELVRYGWGLKLFKKVYTIGEARIRLN CIERLIHTNLLIEVDDVR
 CIKMHDLVRAFVLD MYSKVEHASIVNHGNTLEWHVDNMHNSCKRLSLTCKGMSK
 FPTDLKFPNLSILKLMHEDISLRFPKNFYEE MEKLEVSYDKMKYPLLPSSPQCSVNL
 CVFHLHKCSLVMFDCSCIGNLSNLEVLSFADSAIDLLPSTIGILKKLRLLDLTNCYGL
 CIANGVFKKLVKLEELYMTVVNGGVRKAISL

15

RG2O polynucleotide sequence (SEQ ID NO:118)

TTGTAAAACGACGGCCAGTCGAATCGAACCGTTCGTACGAGAACATCGCTG
 TCCTCTCCTTCATTTGAATCATGATATTGAATATCGATACTTTGACTG
 TAGCTTTGGTCGATTTTAGCAAGATA CATAACTGCCAAACCCATT
 20 GGCTATTTAGCCAAAATATGAAATGGACTGGATTGTTTTCTTC
 TAACACGCACACATCTGGCGATCAGTATCACTCCATTATGAAGACCTAGT
 CAAATTCA TTAACTTCAGTCGTTCTCAAAGTTCAAAGTTCCA ACTT
 CCAACTCCCTCTTTTTCTTCCTCGATTCTGATTGAATCCGAT
 TCTCGACGAAGGAGAGCTGGTCAGAGGGCTGTGATTCTGAGTCTGA
 25 CCTCCGAATCTAGCTGGATTATTCGACACACCAGACCGTACAGGT
 TGCTCATCCC GAAATACTGCTTGCAAACACTGTTGTATCATGCC TAGGAA
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 CTCATTATAAGCTGATCCATATTACATATCTTGAAGAATAATAGGT
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 30 TCCATTTTGTAAGGGATCTGGTTAGTTAAAGGATTGCTACAAC
 AGTATCCCACAAACGATCTATTCCCATTNACTCATCCGCTCAAGATCT
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 TGACAA GTTGGTTACTATGTAGGTACTTCCACAATTAGAATTCCA
 35 GCAATGGATGTTGTTAATGCCATTCTAAACCAGTTGCCAGACACTTAT
 GGAACCTGTTAAGAAACATCTAGGCTACATCATTCCAGCACAAACATG
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 40 AAGCCCTCCTAGTGTACCGCTTGTGAGTCTCAAGATCAAACAT
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 AAGACAAACACTTTGATCACCTGGACTGATCATCCCATTCTGGAA

AAGTTGATTCCATGAAGGCATCGATGCCACAGCATCAACCGATTACAAT
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ACCAAACAACGCTCCCACATGATAGCGTTATGTGGATGGGTGGAGTGG
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5 ATGTTCAGTTATGGTGGAGGCAGTTAGGGAAAAGACGGACCCAAT
TGCTATTCAACAAGCTGTAGCGGATTACCTCGTATAGAGTTAAAAGAAA
GCACTAACCAACAGCAAGAGCTGATAAGCTTCGTGAATGGTCAAGGCCAAC
TCTGGAGAAGGTAAGAATAAATTCTGTAAATACTTGATGACGTCTGGCA
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10 TCGACTTCAAGGTCTTATTGACTTCACGAGACGAACATGTTGCACAGTA
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15 GAAGGATGCACTTCGGTATAGAGCACTATGACCTTCGCAATGTTGCGC
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20 GTGCAGACAAATTGTTAATTGAAAGTGTGATGTTGGGTGTCAAGAT
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25 TGCATGGAGATAAGTCGCTAACGATTCCACAAGACTTTATGAAGGAATG
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CTTGTCTCCTCAATGCTCCACCAACCTCGAGTGCTCATCTCCATGAAT
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30 CGGAAATTAAAGAACGTAAGGTTACTTGATTAAACAGATTGTCATGGTC
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35 TTTGGAAACTTAAACGATTCAAGATCTCAATGGGATGCACTTATATGG
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40 TTCTAAGAGTCTTGTGTTCCAAGTGTGAGTTGAGATACCTTTC
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AAGCTTCGGTTGTGCCAAATGTCAACAAACTGAGCTACCACA
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 5 TATATCTATATGTCTATAATTGATTATATGATGTATTAGTGGATG
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 10 ATCTTGAAGAGCTGAAGTCAAGAAATGTGGTCCATTGAATCGTATT
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 15 CACACCTACCACCAATTAAATATGGGGCACTTGGAGATATCAA
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 20 CTTCTAATAATTATCTGAAGGTGAAAGATCCAACACTTCTAATTGTT
 AACAAATTCAATCTGCAAATGTTCTAAAAATTAAATTACCTGAAA
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 CATTCTGTTGGATATATGTACAGACTGATATTGTCAGAGGAAG
 25 TGAATTACAAGAAGTCACTGATACTATTCTAATGTTGATTCACTCG
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 ATTGGTAACAACATACCATAACAAACAACAACAACCTATATT
 CCCAACCTTGAGGAATTATCTATATTATGACACATGAGTCATGT
 30 ATGGAAGTGCAACAACTGGAATAAATTTCACAACAATCAGAACCCCC
 TCCACAACCTCACAACCATAACACATGTCCGATTGCAAAAGCATTA
 TTGTTTCACCTCTCATGGCAGAACTCTTCAACCTAAAGAGAAATCAA
 TATTGACGAGTGTGATGGTATTGAAGAAATTGTTCAAAAGAGATGATG
 TGGATGAAGAA

35

RG2O deduced polypeptide sequence (SEQ ID NO:119)

MDVNAILKPVAEI LMEPVKKHLGYIISSTKHVRDMSNMR
 ELNAARHAEEDHLD
 RNIRTRLEISNQVRSWLEEVEKIDAKVKALPSDV
 TACCSLKI
 KHEVGREALKLIVEIE
 SATRQHSLITWTDHPIPLGKVDSMKASM
 STASTDYNDFQSREKTFTQALKALEPNN
 40 ASHMIALCGMGGVGKTTMMQRLKKVAKQNR
 MFSYMVEAVIGEK
 TDPIAIQQAVA
 DYLRIELKESTKPARADKLREWFKANS
 GE
 GKNFLVILD
 DVWQSVDLEDIGLSPFP
 NQGVDFKVLLTSRDEHVCTVMGVGSNSILNVGLLIEAEAQSLFQQF
 VETSEPELHKI

GEDIVRKCCGLPIAIKTMACTLRNKRDAWKDALSRIEHYDLRNVAPKVFETSYHN
 LHDKETKS VFLMCGLFPEDFNIPEELMRYGWGLKIFDRVYTFIARNRINTCIEL
 VQTNLLIESDDVGCVKMHDLVRAFVLGMYSEVEHASVNVHGNIPGWTENDPTDSC
 KAISLTCESMSGNIPGDFKFPNLTILKLMHGDKSLRFPQDFYEGMEKLQVISYDKMK
 5 YPMLPLSPQCSTNLRLHLHECSLKMFDSCSIGNMANVEVLSFANSGIEMPLSTIGN
 LKKLRLLDLTDCHGLHITHGVFNNLVKLEELYMGFSDRPDQTRGNISM TDVSYNE
 LAERSKGSALEFQFFENNAQPNNMSFGKLRFKISMGCTLYGGSDYFKKYAVQ
 NTLKLVTNKGEELDSRMNELFVETEMLCLSVDMMNDLGDVCKSSRSPQPSVFKIL
 10 RVFVVSKCVELRYLFTIGVAKDSLNEHLEVDSCNNMEQLICIENAGKETITFLKLKI
 LSLSGLPKLSGLCQNVNKLELPQLIELKLKGIPGFTCIYPQNKL ETSSLLKEEVVIPKL
 ETLQIDEMENLKEIWHYKVNGERVKLRKIEVSNCDKLVNLFPNPMSSLHHLEEL
 EVKKCGSIESLFNIDLDCVDAIGEEDNMRSRLNIKVNSWKLREVWCIGENNSCP
 VSGFQAVESISIESCKRFRNVFTPTTNTNFNMALLEISIDDCGEYMEMENEKSEKSSSEQ
 15 EQTDILSEEVKLQEVTDTISNVVFTSCLIHSFYNNLRKLNLEKYGGVEVVFEIESSTS
 RELVTTYHKQQQQQQPIFPNLEELYLYMDNMMSHVWKCNNWNKFLQQSESPFH
 LTTIHMSDCKSIKYLFSPLMAELLSNLKRINIDECDGI

RG2P polynucleotide sequence (SEQ ID NO:120)

CCCATTGCTATTCAAGGAAGCAGTAGCAGATTACCTCNGTATAGAGCTAA
 20 AGAAAAAAACTAAATCNGCAAGAGCTGATATGCTTCGAAAATGTTAGTTG
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 25 GATGAAGAAGCACAAAGTTGTTCATGGAGTTGTAACAAATTGAGTGA
 TGGTGAATCCCAAGCTTCATAAGATAGGAGAAGATATTGTAAGAAAGTGT
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 30 ACCAGGAGACTAAATATCTTTGCTTGATTGGATTGTTCCGAAGAC
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 GGCTGAAAATGATGTGAGTGGCTTGCCAAAGAATTCAACATGC
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 40 CCGTTCTCCCTCGTCTCCTCAATTGCTCCACCAACCTCGAGTTCT
 TCATCTCCATCAATGCTCATGATGTTGATTGCTCTGTATTGGAAATC
 TGTTAATCTGGAAGTGTGAGCTTGCTAATTCTGGCATTGAATGGTTA

CCTTCCAGAATTGGAAATTGAAGAACGTAAGGCTACTAGATTGACAGA
 TTGTTTGCTTCGTATAGATAAGGGTGTCTAAAAAATTGGTCAAAC
 TTGAAGAGGTTATATGAGAGTTGCTGTTGAAGCAAAAAGCCGGAAAT
 AGAAAAGCCATTAGCTTCACAGATGATACTGCAATGAGATGGCAGAGCG
 5 TTC

RG2P deduced polypeptide sequence (SEQ ID NO:121)

PIAQEAVADYL?IELKEKTKSARADMLRKMLVAKSDGGKNKFLVILDDVVQFVDL
 EDIGLSPLPNQGVNFVLLTSRDVDVCTMMGVEANSILNMKILLDEEAQSLFMEFV
 10 QISSDVPDKLHKIGEDIVRKCCGLPIAKTMALTRNKS KDAWSDALSRLEHHDLHN
 FVNEVFGISYDYLQDQETKYIFLLCGLFPEDYNIPPEELMRYGWGLNLFKKVYTIRE
 ARARLNTCIERLIHTNLLMEGDVVGCVKMHD LALAFVMDMFSKVQDASIVNHGS
 MSGWPENDVSGSCQRISLTCKGMGFPIDLNP NTLIKLHMHD KFLKFPPDFYEQ
 MEKLQVVSFHEMKYPFLPSSPQYCSTNRLV LHLHQCSLMFDCSCSIGNLFNLEVLSF
 15 ANSGIEWLPSRIGNLKKLRLLDLDCFGLRIDKGVLK NLVKLEEYMRVA VRSKKA
 GNRKAISFTDDNCNEMAERS

RG2Q polynucleotide sequence (SEQ ID NO:122)

TGGGGAAGACACAGT GATA GAAA AAAAAAGAAT GTTGTGGAAAAGAGGA
 20 AAATGTTGATTATGCTGTTGTGGCGGT TATAGGGAAAAGACGGACCCT
 ATTGCTCTTCAGAAA ACTGTTGCGGATTACTTGCA TATTGAGCTAAATGA
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 ACTCGGGATGGAGGTAAGAAAAAGTTCTCGTAATACTCGACGATGTTGG
 CAATCTGTTGATTGGAAGATATTGGTTAAGTACTCCTTTCAAATCA
 25 AGGTGTCAACTCAAGGTTGTTGACATCACGAAAGAGAGAAATTGCA
 CAATGATGGGAGTTGAAGCTGATTAAATTCTCAATGTCAAAGTCTTAGAA
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 GTTACCCATTGCCATTAAAACCATGGCTCTTACTTAAGAAATAAAAGA
 30 AAGGATTCATGGAAGGACGC ACTCTCGTTAGAGGACCATGACACTGA
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 GATATT CCTACTGAGGAGTTGGTGAGGTATGGATGGGGCTTAAATCTATT
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 35 TTGAGCGACTCTTGGATTCAAATTGTTGATTGAAAGTAACGATATT CGG
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 40 GAAACTTATGCANGGAGATAAGTCTCAAGGTTCTCAAGACTTTATC
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 TGAATTGGAAGTCCTCAGCTTTTAATTCTAACATTGAATGGTTACCT
 TCCACAATCAGAAATTAAAAAGCTAAGGCTACTAGATTGAGATATTG
 TGATCGTCTCGTATAGAACAAAGGTGTCTGAAAAATTGGTCAAACITG
 5 AAGAACTTTATACTGGATATACATCAGCGTTACAGA

RG2Q deduced polypeptide sequence (SEQ ID NO:123)

GEDTVIEKKKNVVEKRKMFYAVVAVIGEKTDPIALQKTVADYLHIELNESTKLAR
 ADKLCKWFKDNDGGKKFLVILDDVWQSVDLEDIGLSTPFPNQGVNFKVLLSR
 10 KREICTMMGVVEADLILNVKYLEEEEAQKLFHQFVEIGDQYHELHQIGVHVKKCYG
 LPIAIKTMALTLRNKRKDWSKDALSRLEDHDTENVANAVFEMNYRNLQDEETKAI
 FLLCGLFPEDFDIPEELVRYGWGLNFLKKVYTIRKARTRSHTCIERLLDSNLLIESN
 DIRCVKIHDLVRAFVLDMYCKVEHASIVNHGNMRTEYNMADSCKTISLYKSMMSG
 15 FEFPGDLKFPNLTVLKLMDKSLRFPQDFYQSMEKLRVISYDKMKYPLLPSPPQCS
 TNIRVLRLHECSLRMFDCSCIGKLLNLEVLSSFFNSNIEWLPSTIRNLKKLRLLDLRYC
 DRLRIEQGVLKLNVKLEELYTGYSAFTE

RG2S polynucleotide sequence (SEQ ID NO:124)

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 20 CTTATTGATTCTTGTGTTCATGGAGTTGATTTCATTATTACTACCTT
 ACAATTGCTCAGTGATAGATTCCATTAAATTGCTAATTGGTTGCTTC
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 AACGGGGATTGCTGGTGCATTATAACCCATTGCTCAGAGGGCCTGG
 25 TTCCCGTTACAGACCATGTAGGCTACATGATTCTGCAGAAAATATGTG
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 AAATTAAAGGATTGGTGGACCAAGTAGAGAAGGGATCAGAGCAAATGTGGAA
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 GCTTGGACAGAAAGCCTCAAGATAACTGAGCAGATTGAAAGTCTAACAA
 30 GACAGCTCTCCCTGATCAGTGGACTGATGATCCAGTTCTCTAGGAAGA
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 AACAAATTCCACATGGTAGCCTGTGGATGGGTGGAGTAGGAAAGACT
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10 ATTGTCAACCAGGTAATATGCCGAGTGGACTGAAAATGATATAACTGA
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 15 WTCATGATGATGTGAATCTCTAATACCCCATTCAATTGTTGGATGAATG
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RG2S deduced polypeptide sequence (SEQ ID NO:125)

20 MSDPTGIAGAIINPIAQRALVPVTDHVGYMISCRKYVRVMQTKMTELNTSRISVEEH
 ISRNTRNHLQIPSIQKDWLQVEGIRANVENFPIDVITCCSLRIRHKLQKAFKITEQI
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 25 VDFKVLLTSRDSQVCTMMGVEANSIINVGLTEAEAQSLFQQFVETSEPELQKIGED
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 35 LPHLVDLILKGIPGFTVIYPQNKLRTSSLKEEVVIPKLETQLQIDDMENLEEIWPCELS
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 MLPNLKTLKIYMC_{GG}LEHIFTFSALESLTQLQELKIVGCYGMKVIVKKEEDEYGEQ
 QT_{TTTTT}KGASSSSSSSKVVF_{PRLKSIELF}NLPELVGFFLG_{MNEFR}LPSLEEVT
 IKYCSKMMVFAAGGSTAPQLKYI_{HTRLGHTLDQESGLNFHQTSFQSLYGDTS}GPA
 5 TSEGTTWSFHNLIELDMELNYDVKKIIPSSELLQLQKLEKIHVSSCYWVEEVFETAL
 EAAGRNGNSIGFD_{ESSQT}TTTLFNLRLN_{LREM}KLHFLRGLRYIWKS_{NQWT}AF_EF
 PNLTRVHISRCRRLEHVF_{TSSMVGSLLQLQELD}ISWCNHMEEVIVKDADVSVEEDK
 ERESDGKTNKEILVL_{PLR}LSLKLKCLPLKG_{FSLGKEDFSP}LLDTLEIYKC_{PAIT}TFT
 KGNSATPQLKEIETRFGSFYAGEDINSSIIKRSNNRSSNKT_{LINV}K.ILK

10

RG2T polynucleotide sequence (SEQ ID NO:126)

GGAAGACGACAATGGTGCAACGGTTGAAGAAGGTTGTGAAAGATAAGAAG
 ATGTTCCATTATATTGTCGAGGTGGTTGTAGGGGCAAACACTGACCCCAT
 TGCTATCCAGGATACTGTTGCAGATTACCTCAGCATAGAACTGAAAGGAA
 15 ATACGAGAGATGCAAGGGCTTATAAGCTTCGTGAATGCTTAAGGCCCTC
 TCTGGTGGAGGTAAGATGAAGTCCCTAGTAATTCTTGACGATGTATGGAG
 CCTCTGTTGATCTGGATGATATCGGTTAACGTTCTTGCCAAATCAAGGTG
 TTGACTTCAGGTCTTGCTGACATCACGAAACAGTGAATCTGCATGATG
 ATGGGAGCTAGTTAACCTCAATATGTTAACAGACGAGGAAGC
 20 ACATAATTTCGTCGATACGCAGAAATTCTTATGATGCTGATCCCG
 AGCTTATTAAGATAGGAGAACGCTATTGAGAGAAATGTGGTGGTTACCC
 ATTGCCATCAAAACTATGCCGTTACTCTTAGAAATAACGCAAAGATGC
 ATGGAAAGATGCACTTCTCGTTAGAGCACCGTGACACTCATAATGTTG
 TGGCTGATGTTCTAAATTGAGCTACAGCAATATCCAAGACGAGGAGACT
 25 CGGTCGATTTTGCTATGTGGTTGTTCTGAAGACTTGTATATTCC
 TACCGAAGACTTAGTGAGGTATGGATGGGATTGAAAATATTACAGAG
 TGTATACTATGAGACATGCAAGAAAAAGGTTGGACACGTGCATTGAGCGG
 CTTATGCATGCCAACATGTTGATAAAAAGTGATAATGTTGGATTGTCAA
 GATGCATGATCTGGTTCTGCTTTGTTGGCATGTTATCTGAAGTCG
 30 AGCATGCATCAATTGTCACCAGGGATATGCCAGGGTGGTTGAAACT
 GCAAATGATAAGAACAGCTTGTGCAAAAGAATTCTTACATGCAAAGG
 TATGTCGCAATTCTGAAGACCTCACGTTCCAACCTCTCGATCCTGA
 AATTAAATGGATGGAGACGAGTCACTGAGGTTCTGAAGGCTTTATGGA
 GAAATGGAAAACCTCAGGTTATATGATAACATGAAGCAGCCATT
 35 TCTTCCACAATCACTCAATGCTCCAATGTTGAGTGCTTCATCTCCATC
 ACTGCTCATTAATGTTGATTGCTCTCTATTGAAATCTTTGAATCTC
 GAGGTGCTCAGCATTGCTAATTGCCATTAAATTGTTACCCCTCACTAT
 TGGAGATCTGAAGAACGTAAGGCTCTGGATTGACAAATTGTGGTC
 TCTGTATAGCTAATGGCGTCTTAGAAATTGGTCAAACCTGAAGAGCTT
 40 TATATGAGAGTTGATGATCGAGATTGTTTGAAAGCTGATGACAG
 CAAGACCATTACCT

RG2T deduced polypeptide sequence (SEQ ID NO:127)

KTTMVQRLKKVVKDKMFHYIVEVVVGANTDPIAQDTVADYLSIELKGNTNDAR
 AYKLRECFKALSGGGKMKFLVILDDVVSPVLDL DIGLSSLPNQGVDFKVLLTSRNS
 DICMMMGASLIFNLNMLTDEEAHNFFRRYAEISYDADPELIKIGEAIKECGGLPIAI
 5 KTMAVTLRNKRKDAWKDALSRLEHRDTHNVVADVLKLSYSNIQDEETRSIFLLCG
 LFPEDFDIPTEDLVRYGWGLKIFTRVYTMRHARKRLDTIERLMHANMLIKSDNVG
 FVKMHDLVRAFVLGMLSEVEHASIVNHGDMPGFETANDKNSLCKRISLTCKGMS
 AIPEDLTFPNLSILKLMMDGESLRFPEGFYGEMENLQVISYDNMKQPFLPQSLQCSN
 10 VRVLHLHHCSLMFDCCSIGNLLNLEVLSIANSAIKLLPSTIGDLKKLRLLDLTCVGL
 CIANGVFRNLVKLEELYMRVDDRSFFVKADDSTIT

RG2U polynucleotide sequence (SEQ ID NO:128)

GCCTTGTGGATGGGTGGAGTGGAAAGACCACTGTGATGAAGAAGCT
 GAAGGAGGTTGTGGTAGGAAAGAAACTGTTAACATTATGTTGAGGCAG
 15 TTATAGGGAAAAGACAGACCCCCATTGCTATTCAACAAGCTGTTGCCAG
 TACCTTGGTATAAGTCTAACCGAAACCACTAAACCAGCAAGAACTGATAA
 GCTCCGTACATGGTTGCAAACAACACTCAAATGGAGGAAAGAAGAAGTTCC
 TGGAATAACTAGACGATGTATGGCAACCAGTTGATTTGGAAGATATTGGT
 TTAAGTCGTTTCCAATCAAGATGTTGACTTCAAGGTCTGATTACATC
 20 ACGGGACCAATCAGTTGCACTGAGATGGGAGTTAACAGCTGATTAGTTC
 TCAAGGTGAGTGTCCCTGGAGGAAGCGGAAGCACACAGTTGTTCCCTCAA
 TTTTAGAACCTCTGATGATGTCGATCCTGAGCTCAATAAAATCGGAGA
 AGAAATTGTAAGAAGTGTGAGACTACCCATTGCTATCAAACCATGG
 25 CCTGAACCTTAGAAGTAAAAGTAAGGATACATGGAAGAATGCCTTCT
 CGTTTACAACACCATGACATTAACACAATTGGCTACTGTTTCAAAC
 TAGCTATGACAATCTGAAGACGAGGTGACTAAAGCTACTTTTGCTTT
 GTGGTTTATTCCGGAGGACTTCATATTCTACCGAGGACCTATTGAGG
 TATGGATGGGATTGAAGTTATTCAAGGAAGTAGATACTATACGAGAAC
 30 AAGATCCAAGTTGAAAGCCTGCATTGAGCGGCTCATGCATACCAATTGT
 TGATCGAAGGTGATGATGTTAGGTACGTTAAGATGCATGATCTGGTGCCT
 GCCTTGTGATGTTCTAAAGCCGAGCATGCATCTATTGTCAA
 CCATGGTAGTAGTAAGCCAAGGTGGCCTGAAACTGAAAGTGATGTGAGCT
 CCTCTGCAAAGAATTCAACATGCAAGGGTNTG

35 RG2U deduced polypeptide sequence (SEQ ID NO:129)

ALCGMGGVGKTTVMKKLKEVVVGKLFNHYVEAVIGEKTDPPIAQQAVALYLGS
 LTETTKPARTDKLRTWFANNSNGGKKFLVILDDVVWQPVDLEDIGLSRFPNQDVD
 FKVLITSRDQSVCTEMGVKADLVLKVSVLEEAEAHSLFLQFLEPSDDVDPELNKIGE
 EIVKKCCRLPIAKTMA.TLRSKSKDWTKNALSRLQHHDINTIASTVFQTSYDNLEDE
 40 VTKATFLLCGLFPEDFNIPTEDLLRYGWGLKLFKEVDTIREARSKLACIERLMHTN

LLIEGDDVRYVKMHDLVRAFVLDMFSKAEHASIVNHGSSKPRWPETESDVSSCKR-
ISLTCKG?

RG2V polynucleotide sequence (SEQ ID NO:130)

5 CTGTGGAAGACACGAATGATSAAGAAGCTGAAGGAGGTCGTGGAACAAAA
 GAAAATGTTCAATATTATTGTTCAAGTGGTCATAGGAGAGAAAGACAAACC
 CTATTGCTATTCAAGCTGAGCTGATAAGCTTCGTNAATGGTTCGAGGA
 GAAAACACTAAAGAACAGAGCTGATAAGCTTCGTNAATGGTTCGAGGA
 CGATGGAGGAAAGAATAAGTTCCCTGTAATACTTGATGATGTATGGCAGT
10 TTGTCGATCTGAAGATATTGGTTAAGTCCTCTGCCAAATAAAGGTGTC
 AACTTCAAGGTCTTGTGACGTTAAGAGATTACATGTTGCACTCTGAT
 GGGAGCTGAAGCCAATTCAATTCTCAATATAAAAGTTTAAAAGATGTTN
 AAGGACAAAGTTGTCGCCAGTTGCTAAAAATGCAGGTGATGATGAC
 CTGGATCCTGCTTCATGGGATAGCAGATAGTATTGCAAGTAGATGTCA
15 AGGTTGCCCATTGCCATAAAACCATTGCCTTAAGTCTTAAAGGTAGAA
 GCAAGCCTGCGTGGGACCATGCGCTTCTCGTTGGAGAACCATAAAGATT
 GGTAGTGAAGAAGTTGCGTGAAGTTTAAAATTAGCTATGACAATCT
 CCAAGATGAGGTTACTAAATCTATTTTWTACTTGTGCTTATTCCTG
 AAGATTITGATATTCTATTGAGGAGTTGGTGAGGTATGGTGGGCTTG
20 AAATTATTATAGAACAAAACATAAGAGAACAGAAACAGGCTCAA
 CACCTGCACTGAGCGGCTAGGGAGACAAATTGTTATTGGAAGTGATG
 ACATTGGATGCGTCAAGATGCACGATGTGGTGCCTGATTGTTGGTAT
 ATATTCTCAGAAGTCCAGCACGCTTCATTGCAACCAGGTAATGTGTC
 AGAGTGGCTAGAGGAAAATCATAGCATCTACTCTGTAAAAGAATTTCAT
25 TAACATGCAAGGGTATGTCGAGTTCCCAAAGACCTCAAATTCCAAAC
 CTTCAATTGAAACTTATGCATGGAGATAAGTCGNTGAGCTTCCTGA
 AGACTTTATGGAAAGATGGAAAAGGTTCAGGTAAATCATATGATAAT
 TGATGTATCCATTGCTCCCTCATCACTGAATGCTCCACTAACGTTCGA
 GTGCTTCATCTCATTGTCATTAGGATGTTGATTGCTCTCAAT
30 TGGTAATCTCTCAACATGGAAGTGCAGCTTCTAATTCTAACATTG
 AATGGTTACCATCTACAATTGGAAATTGAAAGAAGCTAAGGCTACTAGAT
 TTGACAAATTGTAAGGTCTCGTATAGATAATGGTGTCTAAAAAATT
 GGTCAAACCTGAAGAGCTTATATGGGTGTTAATGTCCGTATGGACAGG
 CCGT

35

RG2V deduced polypeptide sequence (SEQ ID NO:131)

LWKTRM?KKLKEVVEQKKMFNIIVQVVIGEKTNPIAIQQAVADYLSIELKENTKEAR
ADKLR?WFEDDGGKNKFLVILDDVWQFVDLEDIGLSPLPNKGVNFKVLLLRDSH
VCTLMGAEANSILNIKVLKD?GQSLFRQFAKNAGDDLDPAFNGIADSIASRCQGL
40 PIAIKTIALSLKGRSKPAWDHALSRLENHKIGSEEVREVFKISYDNLQDEVTKSIF?L
 CALFPEDFDIPIEELVRYGWGLKLIEAKTIAREARNRLNTCTERLRETNLLFGSDDIG

CVKMHDVVRDFVWYIFSEVQHASIVNHGVSEWLEENHSIYSCKRISLTCKGMSEF
 PKDLKFPNLSILKLMHGDKS?SFPEDFYGKMEKVQVISYDKLMPPLPSSLECSTNV
 RVLHLHYCSLRMFDCSSIGNLLNEVLSFANSNIEWLPSTIGNLKKRLLDLTNCKG
 LRIDNGVLKNLVKLEELYMGVNVRMDQAV

5

RG2W polynucleotide sequence (SEQ ID NO:132)

TTGGGAAAGAGACAATGATGAAGAATTGAAAGAGGTTGTGGTTGAAAAGA
 AAATGTTAACATTATGTGGAGGCAGTTATAGGGGAGAAGACGGACCCC
 ATTGCTATTCAAGCAAGCCGTTGCAGAGTACCTGGTATAATTCTAACAGA
 AACCACTAAGGCAGCAAGAACCGATAAGCTACGTGCATGGCTTCTGACA
 ATTCAGATGGAGGAAGAAAGAAGTCCCTAGTAATACTAGACGATGTATGG
 CATCCGGTTGATATGGAAGATATTGGTTAAGTCGTTCCCAAATCAAGG
 TGTGACTTCAAGGTCTGATTACATCACGGGACCAAGCTGTTGCACTG
 AGATGGGAGTTAAAGCTGATTCAAGGTGAGTGTCTAGAGGAA
 GCTGAAGCACAAGCTTATTCTGCCAACCTTGGAACCTTCTGATGATGT
 CGATCCTGAGCTCCATCAGATTGGAGAAGAAATTGTAAGGAAGTGTGTTG
 GTTACCCATTGCAATAAAACCATGGCCTGCACTCTAGAAGTAAAAGC
 AAGGATACATGGAAGAATGCACTTCTCGTTACAACACCATGACATTAA
 CACAGTCGCGCCTACTGTTTCAAACCAAGCTATGACAATCTCCAAGATG
 20 AGGTGACTGGAGATACTTTGCTATGTGGTTGTTCCGGAGGACTTC
 GATATTCCACTGAAGACTTATTGAAGTATGGATGGGGCTTAAAATTATT
 CAAGGGAGTGGATTCTGTAAGAGAAGCAAGATAACCAGTTGAACGCCTGCA
 TTGAGCGGCTCGTGCATACCAATTGTTGATTGAAAGTGTGTTGGG
 TGCGTCAAGTTGACGATCTGGTGCCTTTATTGTTGATGTTGG
 25 TAAAGCGGAGCATGCTTCGATTGTCACCATGGTAGTAGTAAGCCTGGGT
 GCCCTGAAACTGAAAATGATGTGATCAGGACCTCCTGCAAAAGAATCTCA
 TTAACATGCAAGGGTATGATTGAGTTCTAGTGACCTCAAGTTCCAAA
 TGTCTGATTAAAACCTATGCACTGGAGATAAGTCGCTAAGGTTT

30

RG2W deduced polypeptide sequence (SEQ ID NO:133)

WERDNDEELKEVVVEKKMFNHYVEAVIGEKTDPIAIQQAVAELYLGIILETTEKAAR
 TDKLRAWLSDNSDGGRKFLVILDDVWHPVDMEDIGLSRFPNQGVDFKVLITSRD
 QAVCTEMGVKADSVIKVSLEEAEAQSLFCQLWEPSDDVDPELHQIGEEIVRKCCG
 LPIAJKTMACTLRSKSKDTWKNALSRLQHHDINTVAPTVFQTSYDNLQDEVGDTF
 35 LLCGLFPEDFDIPTEDLLKYGWGLKLFKGVDVSREARYQLNACIERLVHTNLLIESD
 VVGCVKLHDLVRAFILDMFCKAEHASIVNHGSSKPGWPETENDVIRTSCKRISLTCK
 GMIEFSSDLKFPNVLIKLMHGDKSLRF

35

RG5 polynucleotide sequence (SEQ ID NO:134)

40

GGGGGGGTGGGAAGNCGACTCTAGCCCAGAAGNTCTATAATGACCATAA
 AATAAAAGGAAGCTTAGTAAACAAAGCATGGATCTGTGTTCTCAACAAAT

ATTCTGATATTCAGTTGAAAGAACGCCTCGAACATCGGTGTTGAT
TATAAGCATGATGAAACTGTTGGAGAACTTAGCAGAAGGCTTGCAATAGC
TGTCGAAAATGCAAGTTCTTCTTGTGTTGGATGATATTGGCAACATG
AGGTGTGGACTAATTACTCAGAGCCCCATTAAACACTGCAGCTACAGGA
5 ATAATTCTAGTAACAACACTGTAATGATACAGTGCACGAGCAATTGGGTT
GGAAGATATTGATCGAGTAGAATTGATGTCAGATGAAGTAGGATGGAAAT
TGCTTTGAAGAGTATGAACATTAGCAAAGAAAGTGAAGTAGAAAACCTA
CGAGTTTAGGGTTGACATTGTTGTTGTGGTGGCCTCCCCCTAGC
CTT

10

RG5 deduced polypeptide sequence (SEQ ID NO:135)

GGVGKTLAQK?YNDHKIKGSFSKQAWICVSQQYSDISVLKEVLRNIGVDYKHDET
VGELSRRLAIAVENASFFLVLDIWFHEVWTNLLRAPLNTAATGIILVTTRNDTVA
RAIGVEDIHRVELMSDEVGWKLKSMNISKESEVENLRVLGVDIVRLCGGLPLAL

15

RG7 polynucleotide sequence (SEQ ID NO:136)

GGTGGGGTTGGGAAGACAAACGGGCACAAGGAGGCGACTGCCAATACCC
GACTTTATTGATAGAGATGACGAGTCTTATTTCCTACTACTATAGGGA
GGATATTGGTTGCCGAGACGATTGCGCGAAGGGATTCTATCCTT
20 CTTTTTTCCGCGAAGACTCGTCCGGAGGACGGCTATATTCCCTTA
ATATTAGTCTAGCCCAGTCTAGGCCAACCATGGCGATGCGGTAGACCT
CCCAGAGATAGATACTTGATCTTAGAGGATTCACACGTTCAATGGTGGAA
ACTTAAGGAACCGGCTAAGAGTGAACAAACGGAAAAACCTATTCC
ATAGCCTCATCCGGTCGAGGCATTAAACAATCCATCCAATCCTTTCC
25 TTTGGTCTACTCTAATGATGTCGCCGTTGTTGGAAATATCTCTTAT
ACCGACGATTATGGGGATTGCCACTAGCGTTG

The above examples are provided to illustrate the invention but not to limit its scope. Other variants of the invention will be readily apparent to one of ordinary skill in the art and are encompassed by the appended claims. All publications, patents, and patent applications cited herein are hereby incorporated by reference.

WHAT IS CLAIMED IS:

1. An isolated nucleic acid construct comprising an RG polynucleotide which encodes an RG polypeptide having at least 60% sequence identity to an RG polypeptide from an RG family selected from the group consisting of: an RG1 polypeptide, an RG2 polypeptide, an RG3 polypeptide, an RG4 polypeptide, an RG5 polypeptide, and an RG7 polypeptide.
5
2. The nucleic acid construct of claim 1, wherein the RG polynucleotide encodes an RG polypeptide comprising an leucine rich region (LRR).
10
3. The nucleic acid construct of claim 1, wherein the RG polynucleotide encodes an RG polypeptide comprising a nucleotide binding site (NBS).
15
4. The nucleic acid construct of claim 1, wherein the polynucleotide is a full length gene.
15
5. The nucleic acid construct of claim 1, wherein the further encodes a fusion protein.
20
6. The nucleic acid construct of claim 1, wherein the RG1 polypeptide is encoded by an RG1 polynucleotide sequence.
20
7. The nucleic acid construct of claim 6, wherein the RG1 polypeptide is encoded by a polynucleotide sequence selected from the group consisting of SEQ ID NO:1 (RG1A), SEQ ID NO:2 (RG1B), SEQ ID NO: 3 (RG1C), SEQ ID NO:4 (RG1D), SEQ ID NO:5 (RG1E), SEQ ID NO:6 (RG1F), SEQ ID NO:7 (RG1G), SEQ ID NO:8 (RG1H), SEQ ID NO:9 (RG1I), and SEQ ID NO:10 (RG1J).
25
8. The nucleic acid construct of claim 1, wherein the RG2 polypeptide is encoded by an RG2 polynucleotide sequence.
30
9. The nucleic acid construct of claim 8, wherein the RG2 polypeptide is encoded by a polynucleotide sequence selected from the group consisting of: SEQ ID NO:21 (RG2A);

SEQ ID NO:23 (RG2B); SEQ ID NO:25 (RG2C); SEQ ID NO:27 (RG2D); SEQ ID NO:29 (RG2E); SEQ ID NO:31 (RG2F); SEQ ID NO:33 (RG2G); SEQ ID NO:35 (RG2H); SEQ ID NO:37 (RG2I); SEQ ID NO:39 (RG2J); SEQ ID NO:41 (RG2K); SEQ ID NO:43 (RG2L); SEQ ID NO:45 (RG2M); SEQ ID NO:87 (RG2A); SEQ ID NO:89 (RG2B); SEQ ID NO:91 (RG2C); SEQ ID NO:93 (RG2D) and SEQ ID NO:94 (RG2D);
5 SEQ ID NO:96 (RG2E); SEQ ID NO:98 (RG2F); SEQ ID NO:100 (RG2G); SEQ ID NO:102 (RG2H); SEQ ID NO:104 (RG2I); SEQ ID NO:106 (RG2J) and SEQ ID NO:107 (RG2J); SEQ ID NO:109 (RG2K) and (SEQ ID NO:110 (RG2K); SEQ ID NO:112 (RG2L);
10 SEQ ID NO:114 (RG2M); SEQ ID NO:116 (RG2N); SEQ ID NO:118 (RG2O); SEQ ID NO:120 (RG2P); SEQ ID NO:122 (RG2Q); SEQ ID NO:124 (RG2S); SEQ ID NO:126 (RG2T); SEQ ID NO:128 (RG2U); SEQ ID NO:130 (RG2V); and, SEQ ID NO:132 (RG2W).

10. The nucleic acid construct of claim 1, wherein the RG3 polypeptide is encoded by
15 an RG3 polynucleotide sequence.

11. The nucleic acid construct of claim 10, wherein the RG3 polypeptide is encoded by
a polynucleotide sequence as set forth in SEQ ID NO:68.

20 12. The nucleic acid construct of claim 1, wherein the RG4 polypeptide is encoded by
an RG4 polynucleotide sequence.

13. The nucleic acid construct of claim 12, wherein the RG4 polypeptide is encoded by
a polynucleotide sequence as set forth in SEQ ID NO:69.

25 14. The nucleic acid construct of claim 1, wherein the RG5 polypeptide is encoded by
an RG5 polynucleotide sequence.

30 15. The nucleic acid construct of claim 14, wherein the RG5 polypeptide is encoded by
a polynucleotide sequence as set forth in SEQ ID NO:134.

16. The nucleic acid construct of claim 1, wherein the RG7 polypeptide is encoded by an RG7 polynucleotide sequence.
17. The nucleic acid construct of claim 16, wherein the RG7 polypeptide is encoded by 5 a polynucleotide sequence as set forth in SEQ ID NO:136.
18. The nucleic acid construct of claim 1, further comprising a promoter operably linked to the RG polynucleotide.
- 10 19. The nucleic acid construct of claim 18, wherein the promoter is a plant promoter.
20. The nucleic acid construct of of claim 19, wherein the plant promoter is a disease resistance promoter.
- 15 21. The nucleic acid construct of claim 19, wherein the plant promoter is a lettuce promoter.
22. The nucleic acid construct of claim 18, wherein the promoter is a constitutive promoter.
- 20 23. The nucleic acid construct of claim 18, wherein the promoter is an inducible promoter.
24. The nucleic acid construct of claim 18, wherein the promoter is a tissue-specific promoter.
- 25 25. A nucleic acid construct comprising a promoter sequence from an RG gene linked to a heterologous polynucleotide.
- 30 26. A transgenic plant comprising a recombinant expression cassette comprising a promoter operably linked to an RG polynucleotide.

27. The transgenic plant of claim 26, wherein the plant promoter is a plant promoter.

28. The transgenic plant of claim 26, wherein the plant promoter is a viral promoter.

5 29. The transgenic plant of claim 26, wherein the plant promoter is a heterologous promoter.

30. The transgenic plant of claim 26, wherein the plant is lettuce.

10 31. The transgenic plant of claim 26, wherein the RG polynucleotide is selected from the group consisting of SEQ ID NO:1 (RG1A), SEQ ID NO:2 (RG1B), SEQ ID NO: 3 (RG1C), SEQ ID NO:4 (RG1D), SEQ ID NO:5 (RG1E), SEQ ID NO:6 (RG1F), SEQ ID NO:7 (RG1G), SEQ ID NO:8 (RG1H), SEQ ID NO:9 (RG1I), and SEQ ID NO:10 (RG1J).

15 32. The transgenic plant of claim 26, wherein the RG polynucleotide is selected from the group consisting of SEQ ID NO:21 (RG2A); SEQ ID NO:23 (RG2B); SEQ ID NO:25 (RG2C); SEQ ID NO:27 (RG2D); SEQ ID NO:29 (RG2E); SEQ ID NO:31 (RG2F); SEQ ID NO:33 (RG2G); SEQ ID NO:35 (RG2H); SEQ ID NO:37 (RG2I); SEQ ID NO:39 (RG2J); SEQ ID NO:41 (RG2K); SEQ ID NO:43 (RG2L); SEQ ID NO:45 (RG2M); SEQ ID NO:87 (RG2A); SEQ ID NO:89 (RG2B); SEQ ID NO:91 (RG2C); SEQ ID NO:93 (RG2D) and SEQ ID NO:94 (RG2D); SEQ ID NO:96 (RG2E); SEQ ID NO:98 (RG2F); SEQ ID NO:100 (RG2G); SEQ ID NO:102 (RG2H); SEQ ID NO:104 (RG2I); SEQ ID NO:106 (RG2J) and SEQ ID NO:107 (RG2J); SEQ ID NO:109 (RG2K) and (SEQ ID NO:110 (RG2K); SEQ ID NO:112 (RG2L); SEQ ID NO:114 (RG2M); SEQ ID NO:116 (RG2N); SEQ ID NO:118 (RG2O); SEQ ID NO:120 (RG2P); SEQ ID NO:122 (RG2Q); SEQ ID NO:124 (RG2S); SEQ ID NO:126 (RG2T); SEQ ID NO:128 (RG2U); SEQ ID NO:130 (RG2V); and, SEQ ID NO:132 (RG2W).

30 33. The transgenic plant of claim 26, wherein the RG polynucleotide is selected from the group consisting of SEQ ID NO:68 (RG3) and SEQ ID NO:69 (RG4).

34. The transgenic plant of claim 26, wherein the RG polynucleotide comprises a sequence as set forth in SEQ ID NO:134 (RG5).

5 35. The transgenic plant of claim 26, wherein the RG polynucleotide comprises a sequence as set forth in SEQ ID NO:136 (RG7).

10 36. The transgenic plant of claim 26, wherein the RG polynucleotide encodes an RG1 polypeptide selected from the group consisting of SEQ ID NO:11 (RG1A), SEQ ID NO:12 (RG1B), SEQ ID NO: 13 (RG1C), SEQ ID NO:14 (RG1D), SEQ ID NO:15 (RG1E), SEQ ID NO:16 (RG1F), SEQ ID NO:17 (RG1G), SEQ ID NO:18 (RG1H), SEQ ID NO:19 (RG1I), and SEQ ID NO:20 (RG1J).

15 37. The transgenic plant of claim 26, wherein the RG polynucleotide encodes an RG2 polypeptide selected from the group consisting of SEQ ID NO:22 and SEQ ID NO:41 (RG2A); SEQ ID NO:24 and SEQ ID NO:42 (RG2B); SEQ ID NO:43 (RG2C); SEQ ID NO:44 (RG2D); SEQ ID NO:45 (RG2E); SEQ ID NO:46 (RG2F); SEQ ID NO:47 (RG2G); SEQ ID NO:48 (RG2H); SEQ ID NO:49 (RG2I); SEQ ID NO:50 (RG2J); SEQ ID NO:51 (RG2K); SEQ ID NO:52 (RG2L); SEQ ID NO:53 (RG2M); SEQ ID NO:88 (RG2A); SEQ ID NO:90 (RG2B); SEQ ID NO:92 (RG2C); SEQ ID NO:95 (RG2D); SEQ ID NO:97 (RG2E); SEQ ID NO:99 (RG2F); SEQ ID NO:101 (RG2G); SEQ ID NO:103 (RG2H); SEQ ID NO:105 (RG2I); SEQ ID NO:108 (RG2J); SEQ ID NO:111 (RG2K); SEQ ID NO:113 (RG2L); SEQ ID NO:115 (RG2M); SEQ ID NO:117 (RG2N); SEQ ID NO:119 (RG2O); SEQ ID NO:121 (RG2P); SEQ ID NO:123 (RG2Q); SEQ ID NO:125 (RG2S); SEQ ID NO:127 (RG2T); SEQ ID NO:129 (RG2U); SEQ ID NO:131 (RG2V); and, SEQ ID NO:133 (RG2W).

25 38. The transgenic plant of claim 26, wherein the RG polynucleotide encodes an RG3 polypeptide with a sequence as set forth by SEQ ID NO:138.

30 39. The transgenic plant of claim 26, wherein the RG polynucleotide encodes an RG4 polypeptide with a sequence as set forth by SEQ ID NO:139.

40. The transgenic plant of claim 26, wherein the RG polynucleotide encodes an RG5 polypeptide with a sequence as set forth by SEQ ID NO:135.

41. A method of enhancing disease resistance in a plant, the method comprising
5 introducing into the plant a recombinant expression cassette comprising a promoter functional in the plant and operably linked to an RG polynucleotide sequence.

42. The method of claim 41, wherein the plant is a lettuce plant.

10 43. The method of claim 41, wherein the RG polynucleotide encodes an RG polypeptide selected from the group consisting of SEQ ID NO:22 and SEQ ID NO:41 (RG2A); SEQ ID NO:24 and SEQ ID NO:42 (RG2B); SEQ ID NO:43 (RG2C); SEQ ID NO:44 (RG2D); SEQ ID NO:45 (RG2E); SEQ ID NO:46 (RG2F); SEQ ID NO:47 (RG2G); SEQ ID NO:48 (RG2H); SEQ ID NO:49 (RG2I); SEQ ID NO:50 (RG2J); SEQ ID NO:51 (RG2K); SEQ ID NO:52 (RG2L); SEQ ID NO:53 (RG2M); SEQ ID NO:88 (RG2A); SEQ ID NO:90 (RG2B); SEQ ID NO:92 (RG2C); SEQ ID NO:95 (RG2D); SEQ ID NO:97 (RG2E); SEQ ID NO:99 (RG2F); SEQ ID NO:101 (RG2G); SEQ ID NO:103 (RG2H); SEQ ID NO:105 (RG2I); SEQ ID NO:108 (RG2J); SEQ ID NO:111 (RG2K); SEQ ID NO:113 (RG2L); SEQ ID NO:115 (RG2M); SEQ ID NO:117 (RG2N); SEQ ID NO:119 (RG2O);
15 20 SEQ ID NO:121 (RG2P); SEQ ID NO:123 (RG2Q); SEQ ID NO:125 (RG2S); SEQ ID NO:127 (RG2T); SEQ ID NO:129 (RG2U); SEQ ID NO:131 (RG2V); and, SEQ ID NO:133 (RG2W).

44. The method of claim 41, wherein the RG polynucleotide encodes an RG polypeptide selected from the group consisting of SEQ ID NO:138 (RG3); SEQ ID NO:139 (RG4); and
25 SEQ ID NO:135 (RG5).

45. The method of claim 41, wherein the promoter is a tissue-specific promoter or a plant disease resistance promoter.

46. The method of claim 41, wherein the promoter is a constitutive promoter or an inducible promoter.

47. A method of detecting RG resistance genes in a nucleic acid sample, the method
5 comprising:

contacting the nucleic acid sample with an RG polynucleotide to form a hybridization complex; and,

wherein the formation of the hybridization complex is used to detect the RG resistance gene in the nucleic acid sample.

10

48. The method of claim 47, wherein the RG polynucleotide is an RG1 polynucleotide.

49. The method of claim 47, wherein the RG polynucleotide is an RG2 polynucleotide.

15

50. The method of claim 47, wherein the RG polynucleotide is an RG3 polynucleotide, an RG4 polynucleotide, an RG5 polynucleotide or an RG7 polynucleotide.

51. The method of claim 47, wherein the RG resistance gene is amplified prior to the step of contacting the nucleic acid sample with the RG polynucleotide.

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52. The method of claim 51, where the RG resistance gene is amplified by the polymerase chain reaction.

53. The method of claim 47, wherein the RG polynucleotide is labeled.

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54. An RG polypeptide having at least 60% sequence identity to a polypeptide selected from the group consisting of: an RG1 polypeptide, an RG2 polypeptide, an RG3 polypeptide, an RG4 polypeptide, an RG5 polypeptide, and an RG7 polypeptide.

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US98/00615

A. CLASSIFICATION OF SUBJECT MATTER

IPC(6) :Please See Extra Sheet.

US CL :435/6, 91.2, 418, 419; 530/350; 536/23.1, 23.6, 24.1; 800/205

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

U.S. : 435/6, 91.2, 418, 419; 530/350; 536/23.1, 23.6, 24.1; 800/205

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

APS, DIALOG

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	PARAN et al. Development of Reliable PCR-Based Markers Linked to Downy Mildew Resistance Genes in Lettuce. Theor. Appl. Genet. 1993. Vol. 85, No. 8, pages 985-993, see entire article.	1-6, 8, 10, 12, 14, 16, 18-30, 41-42, 45-54
Y	KESSELI et al. Analysis of a Detailed Genetic Linkage Map of <i>Lactuca sativa</i> (Lettuce) Constructed From RFLP and RAPD Markers. Genetics. April 1994. Vol. 136, No. 4, pages 1435-1446, see entire document.	1-6, 8, 10, 12, 14, 16, 18-30, 41-42, 45-54
Y	MICHELMORE, RW. Isolation of Disease Resistance Genes from Crop Plants. Current Opinion in Biotechnology. 1995. Vol. 6, No. 2, pages 145-152, see entire document.	1-6, 8, 10, 12, 14, 16, 18-30, 41-42, 45-54

 Further documents are listed in the continuation of Box C. See patent family annex.

Special categories of cited documents:	"T"	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
"A" document defining the general state of the art which is not considered to be of particular relevance	"X"	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
"B" earlier document published on or after the international filing date	"Y"	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art
"L" document which may throw doubt on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"Δ"	document member of the same patent family
"O" document referring to an oral disclosure, use, exhibition or other means		
"P" document published prior to the international filing date but later than the priority date claimed		

Date of the actual completion of the international search

14 MARCH 1998

Date of mailing of the international search report

13 APR 1998

Name and mailing address of the ISA/US
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INTERNATIONAL SEARCH REPORT

International application No. PCT/US98/00615

C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	PARAN et al. Recent Amplification of Triose Phosphate Isomerase Related Sequences in Lettuce. Genome. 1992. Vol. 35, No. 4, pages 627-635, see entire document.	1-6, 8, 10, 12, 14, 16, 18-30, 41-42, 45-54
Y	PARAN et al. Identification of Restriction Fragment Length Polymorphism and Random Amplified Polymorphic DNA markers linked to Downy Mildew Resistance Genes in Lettuce, Using Near-Isogenic Lines. Genome. 1991. Vol. 34, No. 6, pages 1021-1027, see entire document.	1-6, 8, 10, 12, 14, 16, 18-30, 41-42, 45-54

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US98/00615

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:

2. Claims Nos.: 7, 9, 11, 13, 15, 17, 31-40, 43-44 because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:

these claims are drawn to numerous sequences identified by SEQ ID NOs. However, since no computer readable form was submitted, no meaningful search could be carried out.

3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:

4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

The additional search fees were accompanied by the applicant's protest.

No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

International application No.
PCT/US98/00615

A. CLASSIFICATION OF SUBJECT MATTER:

IPC (6):

A01H 1/00; C07H 21/04; C07K 14/00; C12N 5/04, 5/10; C12P 19/34; C12Q 1/68